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(71) Applicant (for all designated States except US): **EX-ELIXIS, INC.** [US/US]; P.O. Box 511, 170 Harbor Way, South San Francisco, CA 94083-0511 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **BUHR, Chris, A.** [US/US]; 355 Belmont Avenue, Redwood City, CA 94061 (US). **BAIK, Tae-Gon** [KR/US]; 1125 Balclutha Drive, #105, Foster City, CA 94404 (US). **MA, Sunghoon** [KR/US]; 1110 Polynesia Drive, #109, Foster City, CA 94404 (US). **TESFAI, Zerom** [ER/US]; 4704 Springlake Drive, San Leandro, CA 94578 (US). **WANG, Longcheng** [CN/US]; 22 Skypark Circle, South San Francisco, CA 94080 (US). **CO, Erick, Wang** [US/US]; 921 Salt Court, Redwood City, Ca 94065 (US). **EPSHTEYN, Sergey** [US/US]; 2922 Southwycke Terrace, Fremont, CA 94536 (US). **KENNEDY, Abigail, R.** [US/US]; 325 West Lowell Avenue, Tracy, CA 95376 (US). **CHEN, Baili** [US/US]; 1054 Colorado Avenue, Palo Alto, CA 94303 (US). **DUBENKO, Larisa** [US/US]; 3326 Balboa Street, San Francisco, CA 94121 (US). **ANAND, Neel Kumar** [GB/US]; 2713 South Norfolk Street, Apt.#206, San Mateo, CA 94403 (US). **TSANG, Tsze, H.** [US/US]; 8367 Kent Drive, El Cerrito, CA 94530 (US). **NUSS, John, M.** [US/US]; 16 Woodranch Circle, Danville, CA 94056 (US). **PETO, Csaba, J.** [HU/US]; 965 Shorepoint Court, #305, Alameda, CA 94501 (US). **RICE, Kenneth, D.** [US/US]; 802 Autumn Lane, Mill Valley, CA 94941 (US). **IBRAHIM, Mohamed, Abdulkader** [US/US]; 3380 Lubich Drive, Mountain View, CA 94040 (US). **SCHNEPP,**

Kevin, Luke [US/US]; 8768 Newry Court, Elk Grove, CA 95624 (US). **SHI, Xian** [CN/US]; 157 Elm Street, Apt. 106, San Mateo, CA 94401 (US). **LEAHY, James, William** [US/US]; 1185 Camellia Court, San Leandro, CA 94577 (US). **CHEN, Jeff** [US/US]; 417 Fulton Street #6, San Francisco, CA 94102 (US). **DALRYMPLE, Lisa, Esther** [US/US]; 387 17th Avenue, San Francisco, CA 94121 (US). **FORSYTH, Timothy, Patrick** [US/US]; 2071A Isabelle Avenue, San Mateo, CA 94403 (US). **HUYNH, Tai, Phat** [US/US]; 1530 6th Avenue, #2, Oakland, CA 94606 (US). **MANN, Grace** [US/US]; 231 Callippe Court, Brisbane, Ca 94005 (US). **MANN, Lary, Wayne** [US/US]; 780 Bair Island Road, Apt. 304, Redwood City, CA 94063 (US). **TAKEUCHI, Craig, Stacy** [CA/US]; 1090 Carolan Avenue, #302, Burlingame, CA 94010 (US).

(74) Agents: **GRIEDEL, Brian** et al.; Exelixis, Inc., P.O. Box 511, 170 Harbor Way, South San Francisco, CA 94083-0511 (US).

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(54) Title: PROTEIN KINASE MODULATORS AND METHODS OF USE

(57) Abstract: This invention relates to compounds for modulating protein kinase enzymatic activity for modulating cellular activities such as proliferation, differentiation, programmed cell death, migration and chemoinvasion, and to pharmaceutical compositions containing such compounds. Even more specifically, the invention relates to compounds that inhibit, regulate and/or modulate kinases, particularly Checkpoint Kinases, even more particularly Checkpoint Kinase 1, or Chk1. Methods of therapeutically or prophylactically using the compounds and compositions to treat kinase-dependent diseases and conditions are also an aspect of the invention, and include methods of treating cancer, as well as other disease states associated with unwanted angiogenesis and/or cellular proliferation, by administering effective amounts of such compounds.



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Protein Kinase Modulators and Methods of Use

BACKGROUND OF THE INVENTION

Field of the Invention

[001] This invention relates to compounds for modulating protein kinase enzymatic activity for modulating cellular activities such as proliferation, differentiation, programmed cell death, migration and chemoinvasion, and to pharmaceutical compositions containing such compounds. Even more specifically, the invention relates to compounds that inhibit, regulate and/or modulate kinases, particularly Checkpoint Kinases, even more particularly Checkpoint Kinase 1, or Chk1. Methods of therapeutically or prophylactically using the compounds and compositions to treat kinase-dependent diseases and conditions are also an aspect of the invention, and include methods of treating cancer, as well as other disease states associated with unwanted angiogenesis and/or cellular proliferation, by administering effective amounts of such compounds.

Summary of Related Art

[002] Cells reproduce by duplicating their contents and then dividing in two. This cell division cycle is the fundamental means by which all living things are propagated. In unicellular species, such as bacteria and yeasts, each cell division produces an additional organism. In multicellular species many rounds of cell division are required to make a new individual, and cell division is needed in the adult body too, to replace cells that are lost by wear and tear or by programmed cell death. Thus an adult human must manufacture many millions of new cells each second simply to maintain the status quo, and if all cell division is halted – for example, by a large dose of ionizing radiation – the individual will die within a few days.

[003] The cell division cycle is the subject of a control system that coordinates the cycle as a whole. A multitude of genes, proteins and other cellular machinery directly or indirectly regulate progression of a cell through the cell division cycle. Regulatory elements can either help stimulate an increase in cell numbers or help to inhibit it. Uncontrolled cell proliferation is the insignia of cancer, and can be manifested by a

deregulation of the cell division cycle in one of two ways - making stimulatory genes hyperactive or inhibitory genes inactive.

- [004] It can be understood, therefore, that agents that modulate the cell cycle, and thus hyperproliferation, could be used to treat various disease states associated with uncontrolled or unwanted cell proliferation. In addition to oncological indications, altered cell cycle division signaling is implicated in numerous other pathological diseases. These include, but are not limited to antiparasitics (See, Gray et al., *Curr. Med. Chem.* 6, 859-875 (1999)); potential antivirals (See, Schang et al., *J. Virol.* 74, 2107-2120 (2000)); cardiovascular maladies such as atherosclerosis or restenosis (See Braun-Dullaeus et al., *Circulation*, 98, 82-89 (1998)); and states of inflammation and immunological disorders, such as arthritis (See, Taniguchi et al., *Nature Med.*, 5, 760-767(1999)) or psoriasis.
- [005] Mechanisms of cell proliferation are under active investigation at cellular and molecular levels. At the cellular level, de-regulation of signaling pathways, loss of cell cycle controls, unbridled angiogenesis or stimulation of inflammatory pathways are under scrutiny, while at the molecular level, these processes are modulated by various proteins, among which protein kinases are prominent suspects. Overall abatement of proliferation may also result from programmed cell death, or apoptosis, which is also regulated via multiple pathways, some involving proteolytic enzyme proteins. Among the candidate regulatory proteins, protein kinases are a family of enzymes that catalyze phosphorylation of proteins, in particular the hydroxyl group of specific tyrosine, serine, or threonine residues in proteins. The consequences of this seemingly simple activity are staggering, and typically, such phosphorylation dramatically perturbs the function of the protein, and thus protein kinases are pivotal in the regulation of a wide variety of cellular processes, including metabolism, cell proliferation, cell differentiation, and cell survival. Of the many different cellular functions in which the activity of protein kinases is known to be required, some processes represent attractive targets for therapeutic intervention for certain disease states. Two examples are cell-cycle control and angiogenesis, in which protein kinases play a pivotal role; these processes are essential for the growth of solid tumors as well as for other diseases.
- [006] CDKs constitute a class of enzymes that play critical roles in regulating the transitions between different phases of the cell cycle, such as the progression from a quiescent stage in G.sub.1 (the gap between mitosis and the onset of DNA replication for a new round of cell division) to S (the period of active DNA synthesis), or the progression

from G.sub.2 to M phase, in which active mitosis and cell-division occur. See, e.g., the articles compiled in *Science*, vol. 274 (1996), pp. 1643-1677; and *Ann. Rev. Cell Dev. Biol.*, vol. 13 (1997), pp. 261-291. CDK complexes are formed through association of a regulatory cyclin subunit (e.g., cyclin A, B1, B2, D1, D2, D3, and E) and a catalytic kinase subunit (e.g., cdc2 (CDK1), CDK2, CDK4, CDK5, and CDK6). As the name implies, the CDKs display an absolute dependence on the cyclin subunit in order to phosphorylate their target substrates, and different kinase/cyclin pairs function to regulate progression through specific portions of the cell cycle.

[007] The D cyclins are sensitive to extracellular growth signals and become activated in response to mitogens during the G.sub.1 phase of the cell cycle. CDK4/cyclin D plays an important role in cell cycle progression by phosphorylating, and thereby inactivating, the retinoblastoma protein (Rb). Hypophosphorylated Rb binds to a family of transcriptional regulators, but upon hyperphosphorylation of Rb by CDK4/cyclin D, these transcription factors are released to activate genes whose products are responsible for S phase progression. Rb phosphorylation and inactivation by CDK4/cyclin D permit passage of the cell beyond the restriction point of the G.sub.1 phase, whereupon sensitivity to extracellular growth or inhibitory signals is lost and the cell is committed to cell division. During late G.sub.1, Rb is also phosphorylated and inactivated by CDK2/cyclin E, and recent evidence indicates that CDK2/cyclin E can also regulate progression into S phase through a parallel pathway that is independent of Rb phosphorylation (see Lukas et al., "Cyclin E-induced S Phase Without Activation of the pRb/E2F Pathway," *Genes and Dev.*, vol. 11 (1997), pp. 1479-1492).

[008] The progression from G.sub.1 to S phase, accomplished by the action of CDK4/cyclin D and CDK2/cyclin E, is subject to a variety of growth regulatory mechanisms, both negative and positive. Growth stimuli, such as mitogens, cause increased synthesis of cyclin D1 and thus increased functional CDK4. By contrast, cell growth can be "reined in," in response to DNA damage or negative growth stimuli, by the induction of endogenous inhibitory proteins. These naturally occurring protein inhibitors include p.sub.21.sup.WAF1/CIP1, p27.sup.KIP1, and the p16.sup.INK4 family, the latter of which inhibit CDK4 exclusively (see Harper, "Cyclin Dependent Kinase Inhibitors," *Cancer Surv.*, vol. 29 (1997), pp. 91-107). Aberrations in this control system, particularly those that affect the function of CDK4 and CDK2, are implicated in the advancement of cells to the highly proliferative state characteristic of malignancies, such as familial

melanomas, esophageal carcinomas, and pancreatic cancers (see, e.g., Hall and Peters, "Genetic Alterations of Cyclins, Cyclin-Dependent Kinases, and CDK Inhibitors in Human Cancer," *Adv. Cancer Res.*, vol. 68 (1996), pp.67-108; and Kamb et al., "A Cell Cycle Regulator Potentially Involved in Genesis of Many Tumor Types," *Science*, vol. 264 (1994), pp. 436-440).

[009] Over-expression of cyclin D1 is linked to esophageal, breast, and squamous cell carcinomas (see, e.g., DelSal et al., "Cell Cycle and Cancer: Critical Events at the G.sub.1 Restriction Point," *Critical Rev. Oncogenesis*, vol. 71 (1996), pp. 127-142). Genes encoding the CDK4-specific inhibitors of the p16 family frequently have deletions and mutations in familial melanoma, gliomas, leukemias, sarcomas, and pancreatic, non-small cell lung, and head and neck carcinomas (see Nobori et al., "Deletions of the Cyclin-Dependent Kinase-4 Inhibitor Gene in Multiple Human Cancers," *Nature*, vol. 368 (1994), pp. 753-756). Amplification and/or overexpression of cyclin E has also been observed in a wide variety of solid tumors, and elevated cyclin E levels have been correlated with poor prognosis. In addition, the cellular levels of the CDK inhibitor p27, which acts as both a substrate and inhibitor of CDK2/cyclin E, are abnormally low in breast, colon, and prostate cancers, and the expression levels of p27 are inversely correlated with the stage of disease (see Loda et al., "Increased Proteasome-dependent Degradation of the Cyclin-Dependent Kinase Inhibitor p27 in Aggressive Colorectal Carcinomas," *Nature Medicine*, vol. 3 (1997), pp. 231-234). Recently there is evidence that CDK4/cyclin D might sequester p27, as reviewed in Sherr, et al., *Genes Dev.*, Vol. 13 (1999), pp. 1501-1512. The p21 proteins also appear to transmit the p53 tumor-suppression signal to the CDKs; thus, the mutation of p53 in approximately 50% of all human cancers may indirectly result in deregulation of CDK activity.

[010] In at least one instance modulation of protein kinase activity as been used as a treatment for an oncological condition. For example, modulation of protein kinase activity for the treatment of chronic myeloid leukemia (CML) and gastrointestinal stroma cancers has been demonstrated successfully with the FDA approval of Gleevec® (imatinib mesylate, produced by Novartis Pharmaceutical Corporation of East Hanover, NJ). Gleevec is a selective Abl kinase inhibitor.

[011] Another series of protein kinases are those that play an important role as a checkpoint in cell cycle progression. See, e.g., Melo et al., *Current Opinion in Cell Biology* 2002, 14:237-245 (2003). Proliferation of eukaryotic cells is controlled by cell

cycle checkpoint pathways, which mediate progression through critical transitions of the cell cycle (reviewed by Walworth (2000), *Curr. Opin. Cell Biol.* 12, 697-704). Checkpoints prevent cell cycle progression at inappropriate times, such as in response to DNA damage, and maintain the metabolic balance of cells while the cell is arrested, and in some instances can induce apoptosis (programmed cell death) when the requirements of the checkpoint have not been met. See, e.g., O'Connor, *Cancer Surveys*, 29, 151-182 (1997); Nurse, *Cell*, 91, 865-867 (1997); Hartwell et al., *Science*, 266, 1821-1828 (1994); Hartwell et al., *Science*, 246, 629-634 (1989). Checkpoint control can occur in the G1 phase (prior to DNA synthesis) and in G2, prior to entry into mitosis.

[012] One series of checkpoints monitors the integrity of the genome and, upon sensing DNA damage, these "DNA damage checkpoints" block cell cycle progression in G.sub.1 & G.sub.2 phases, and slow progression through S phase. O'Connor, *Cancer Surveys*, 29, 151-182 (1997); Hartwell et al., *Science*, 266, 1821-1828 (1994). This action enables DNA repair processes to complete their tasks before replication of the genome and subsequent separation of this genetic material into new daughter cells takes place. Importantly, the most commonly mutated gene in human cancer, the p53 tumor suppressor gene, produces a DNA damage checkpoint protein that blocks cell cycle progression in G.sub.1 phase and/or induces apoptosis (programmed cell death) following DNA damage. Hartwell et al., *Science*, 266, 1821-1828 (1994). The p53 tumor suppressor has also been shown to strengthen the action of a DNA damage checkpoint in G.sub.2 phase of the cell cycle. See, e.g., Bunz et al., *Science*, 28, 1497-1501 (1998); Winters et al., *Oncogene*, 17, 673-684 (1998); Thompson, *Oncogene*, 15, 3025-3035 (1997).

[013] Given the pivotal nature of the p53 tumor suppressor pathway in human cancer, therapeutic interventions that exploit vulnerabilities in p53-defective cancer have been actively sought. One emerging vulnerability lies in the operation of the G.sub.2 checkpoint in p53 defective cancer cells. Cancer cells, because they lack G.sub.1 checkpoint control, are particularly vulnerable to abrogation of the last remaining barrier protecting them from the cancer killing effects of DNA-damaging agents: the G.sub.2 checkpoint. The G.sub.2 checkpoint is regulated by a control system that has been conserved from yeast to humans. Important in this conserved system is a kinase, CHK1, which transduces signals from the DNA-damage sensory complex to inhibit activation of the cyclin B/Cdc2 kinase, which promotes mitotic entry. See, e.g., Peng et al., *Science*, 277, 1501-1505 (1997); Sanchez et al., *Science*, 277, 1497-1501 (1997). Inactivation of

CHK1 has been shown to both abrogate G.sub.2 arrest induced by DNA damage inflicted by either anticancer agents or endogenous DNA damage, as well as result in preferential killing of the resulting checkpoint defective cells. See, e.g., Nurse, *Cell*, 91, 865-867 (1997); Weinert, *Science*, 277, 1450-1451 (1997); Walworth et al., *Nature*, 363, 368-371 (1993); and Al-Khodairy et al., *Molec. Biol. Cell.*, 5, 147-160 (1994).

[014] Selective manipulation of checkpoint control in cancer cells could afford broad utilization in cancer chemotherapeutic and radiotherapy regimens and may, in addition, offer a common hallmark of human cancer "genomic instability" to be exploited as the selective basis for the destruction of cancer cells. A number of factors place CHK1 as a pivotal target in DNA-damage checkpoint control. The elucidation of inhibitors of this and functionally related kinases such as CDS1/CHK2, a kinase recently discovered to cooperate with CHK1 in regulating S phase progression (see Zeng et al., *Nature*, 395, 507-510 (1998); Matsuoka, *Science*, 282, 1893-1897 (1998)), could provide valuable new therapeutic entities for the treatment of cancer.

[015] Another group of kinases are the tyrosine kinases. Tyrosine kinases can be of the receptor type (having extracellular, transmembrane and intracellular domains) or the non-receptor type (being wholly intracellular). At least one of the non-receptor protein tyrosine kinases, namely, LCK, is believed to mediate the transduction in T-cells of a signal from the interaction of a cell-surface protein (Cd4) with a cross-linked anti-Cd4 antibody. A more detailed discussion of non-receptor tyrosine kinases is provided in Bolen, *Oncogene*, 8, 2025-2031 (1993), which is incorporated herein by reference.

[016] Receptor-type tyrosine kinases are comprised of a large number of transmembrane receptors with diverse biological activity. In fact, about 20 different subfamilies of receptor-type tyrosine kinases have been identified. One tyrosine kinase subfamily, designated the HER subfamily, is comprised of EGFR (HER1), HER2, HER3, and HER4. Ligands of this subfamily of receptors identified so far include epithelial growth factor, TGF-alpha, amphiregulin, HB-EGF, betacellulin and heregulin. Another subfamily of these receptor-type tyrosine kinases is the insulin subfamily, which includes INS-R, IGF-IR, and IR-R. The PDGF subfamily includes the PDGF-alpha and beta receptors, CSFIR, c-kit and FLK-II. Then there is the FLK family, which is comprised of the kinase insert domain receptor (KDR), fetal liver kinase-1 (FLK-1), fetal liver kinase-4 (FLK-4) and the fms-like tyrosine kinase-1 (flt-1). The PDGF and FLK families are usually considered together due to the similarities of the two groups. For a detailed discussion of the

receptor-type tyrosine kinases, see Plowman et al., *DN&P* 7(6): 334-339, 1994, which is hereby incorporated by reference.

[017] The non-receptor type of tyrosine kinases is also comprised of numerous subfamilies, including Src, Frk, Btk, Csk, Abl, Zap70, Fes/Fps, Fak, Jak, Ack, and LIMK. Each of these subfamilies is further sub-divided into varying receptors. For example, the Src subfamily is one of the largest and includes Src, Yes, Fyn, Lyn, Lck, Blk, Hck, Fgr, and Yrk. The Src subfamily of enzymes has been linked to oncogenesis. For a more detailed discussion of the non-receptor type of tyrosine kinases, see Bolen, *Oncogene*, 8:2025-2031 (1993), which is hereby incorporated by reference.

[018] In addition to its role in cell-cycle control, protein kinases also play a crucial role in angiogenesis, which is the mechanism by which new capillaries are formed from existing vessels. When required, the vascular system has the potential to generate new capillary networks in order to maintain the proper functioning of tissues and organs. In the adult, however, angiogenesis is fairly limited, occurring only in the process of wound healing and neovascularization of the endometrium during menstruation. See Merenmies, J., Parada, L. F., Henkemeyer, M., *Cell Growth & Differentiation*, 8, 3-10 (1997). On the other hand, unwanted angiogenesis is a hallmark of several diseases, such as retinopathies, psoriasis, rheumatoid arthritis, age-related macular degeneration, and cancer (solid tumors). Folkman, *Nature Med.*, 1, 27-31 (1995). Protein kinases which have been shown to be involved in the angiogenic process include three members of the growth factor receptor tyrosine kinase family: VEGF-R2 (vascular endothelial growth factor receptor 2, also known as KDR (kinase insert domain receptor) and as FLK-1); FGF-R (fibroblast growth factor receptor); and TEK (also known as Tie-2).

[019] VEGF-R2, which is expressed only on endothelial cells, binds the potent angiogenic growth factor VEGF and mediates the subsequent signal transduction through activation of its intracellular kinase activity. Thus, it is expected that direct inhibition of the kinase activity of VEGF-R2 will result in the reduction of angiogenesis even in the presence of exogenous VEGF (see Strawn et al., *Cancer Research*, 56, 3540-3545 (1996)), as has been shown with mutants of VEGF-R2 which fail to mediate signal transduction. Millauer et al., *Cancer Research*, 56, 1615-1620 (1996). Furthermore, VEGF-R2 appears to have no function in the adult beyond that of mediating the angiogenic activity of VEGF. Therefore, a selective inhibitor of the kinase activity of VEGF-R2 would be expected to exhibit little toxicity.

[020] Similarly, FGF-R binds the angiogenic growth factors aFGF and bFGF and mediates subsequent intracellular signal transduction. Recently, it has been suggested that growth factors such as bFGF may play a critical role in inducing angiogenesis in solid tumors that have reached a certain size. Yoshiji et al., *Cancer Research*, 57, 3924-3928 (1997). Unlike VEGF-R2, however, FGF-R is expressed in a number of different cell types throughout the body and may or may not play important roles in other normal physiological processes in the adult. Nonetheless, systemic administration of a small molecule inhibitor of the kinase activity of FGF-R has been reported to block bFGF-induced angiogenesis in mice without apparent toxicity. Mohammad et al., *EMBO Journal*, 17, 5996-5904 (1998).

[021] TEK (also known as Tie-2) is another receptor tyrosine kinase expressed only on endothelial cells which has been shown to play a role in angiogenesis. The binding of the factor angiopoietin-1 results in autophosphorylation of the kinase domain of TEK and results in a signal transduction process which appears to mediate the interaction of endothelial cells with peri-endothelial support cells, thereby facilitating the maturation of newly formed blood vessels. The factor angiopoietin-2, on the other hand, appears to antagonize the action of angiopoietin-1 on TEK and disrupts angiogenesis. Maisonpierre et al., *Science*, 277, 55-60 (1997).

[022] As a result of the above-described developments, it has been proposed to treat angiogenesis by the use of compounds inhibiting the kinase activity of VEGF-R2, FGF-R, and/or TEK. For example, WIPO International Publication No. WO 97/34876 discloses certain cinnoline derivatives that are inhibitors of VEGF-R2, which may be used for the treatment of disease states associated with abnormal angiogenesis and/or increased vascular permeability such as cancer, diabetes, psoriasis, rheumatoid arthritis, Kaposi's sarcoma, haemangioma, acute and chronic nephropathies, atheroma, arterial restinosis, autoimmune diseases, acute inflammation and ocular diseases with retinal vessel proliferation.

[023] In addition to the protein kinases identified above, many other protein kinases have been considered to be therapeutic targets, and numerous publications disclose inhibitors of kinase activity, as reviewed in the following: McMahon et al., *Oncologist*, 5, 3-10 (2000); Garcia-Echeverria et al., *Med. Res. Rev.*, 20, 28-57 (2000); Holash et al., *Oncogene*, 18, 5356-5362 (1999); Stover et al., *Curr. Opin. Drug Disc. Dev.*, 2, 274-285 (1999); Toledo et al., *Curr Med. Chem.*, 6, 775-805 (1999); Thomas et al., *J. Biol. Chem.*,

274, 36684-36692 (1992); Cohen, *Curr. Op. Chem. Biol.*, 10, 544-549 (1999); Adams et al., *Curr. Opin. Drug Disc. Dev.*, 2, 96-109 (1999); McMahon et al., *Curr. Opin. Drug Disc. Dev.*, 1, 131-146 (1998); and Strawn et al., *Exp. Opin. Invest. Drugs*, 7, 553-573 (1998).

[024] It can be understood, therefore, that there is a need, for effective inhibitors of protein kinases in order to treat or prevent disease states associated with abnormal cell proliferation. Moreover, as would be understood by those skilled in the art, it is desirable for kinase inhibitors to possess both high affinity for the target kinase as well as high selectivity versus other protein kinases. Small-molecule compounds that may be readily synthesized and are potent inhibitors of cell proliferation are those, for example, that are inhibitors of one or more protein kinases, such as CHK1, CHK2, VEGF, CDKs or CDK/cyclin complexes and both receptor and non-receptor tyrosine kinases.

SUMMARY OF THE INVENTION

[025] Based on the aforementioned evidence, we contemplated that highly specific inhibitors of protein kinase, such as Chk1 kinase, will have a synergistic cytotoxic effect with DNA damaging agents in many kinds of tumor cells, whilst not causing excessive toxicity in non-tumor cells. Thus, one aspect of the invention is selective protein kinase modulators, particularly inhibitors like Chk1. Methods of the invention include both use of such compounds in a synergistic fashion as mention, and also as single agents for cancer therapy.

[026] We have unexpectedly found that analogs of 3-amino-pyrazine-2-carboxamide and related compounds effectively inhibit protein kinases such as Chk1 kinase. The compounds disclosed herein are useful for inhibiting the protein kinase and, therefore, useful for studying the role of protein kinase in various biological pathways as well as to augment the efficacy of cancer therapeutics, including DNA damaging agents. Accordingly, the invention also comprises pharmaceutical compositions comprising the compounds of the invention, as well as methods of using compounds of the invention to inhibit protein kinase and thereby modulate kinase dependent diseases or conditions.

[027] In another aspect, the invention provides methods of screening for modulators of kinase activity. The methods comprise combining a composition of the invention, a

kinase, and at least one candidate agent and determining the effect of the candidate agent on the kinase activity.

[028] In yet another aspect, the invention also provides pharmaceutical kits comprising one or more containers filled with one or more of the ingredients of pharmaceutical compounds and/or compositions of the present invention, including, one or more kinase enzyme activity modulators as described herein. Such kits can also include, for example, other compounds and/or compositions (e.g., diluents, permeation enhancers, lubricants, and the like), a device(s) for administering the compounds and/or compositions, and written instructions in a form prescribed by a governmental agency regulating the manufacture, use or sale of pharmaceuticals or biological products, which instructions can also reflect approval by the agency of manufacture, use or sale for human administration.

[029] In still yet another aspect, the invention also provides a diagnostic agent comprising a compound of the invention and, optionally, pharmaceutically acceptable adjuvants and excipients.

[030] These and other features and advantages of the present invention will be described in more detail below.

DETAILED DESCRIPTION OF THE INVENTION

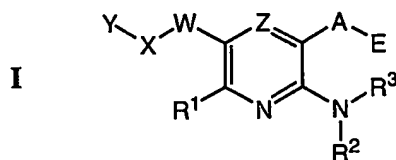
[031] The compounds set forth in the detailed description are useful as pharmaceuticals in mammals, especially humans, for modulating protein kinase enzymatic activity for modulating cellular activities such as uncontrolled, abnormal, or unwanted proliferation, differentiation, programmed cell death, migration and chemoinvasion. It is appreciated, however, that in some cases the cells may not be in a hyper- or hypo- proliferative and/or migratory state (abnormal state) and still require treatment. For example, during wound healing, the cells may be proliferating "normally," but proliferation and migration enhancement may be desired. Alternatively, reduction in "normal" cell proliferation and/or migration rate may be desired. Furthermore, such compounds may be used to prevent de-differentiation of post-mitotic tissue and/or cells.

[032] Diseases or disorders associated with uncontrolled, abnormal, and/or unwanted cellular activities, especially cellular proliferation include, but are not limited to the conditions set forth in the background and to the following:

- [033] a variety of cancers, including, but not limited to carcinoma, hematopoietic tumors of lymphoid lineage, hematopoietic tumors of myeloid lineage, tumors of mesenchymal origin, tumors of the central and peripheral nervous system and other tumors including melanoma, seminoma and Kaposi's sarcoma and the like, and those further discussed below;
- [034] a disease process which features abnormal cellular proliferation, e.g., benign prostatic hyperplasia, familial adenomatosis polyposis, neuro-fibromatosis, pulmonary fibrosis, hypertrophic scar formation, transplantation rejection, nodosum, endotoxic shock, and a variety of cardiovascular indications, including, but not limited to atherosclerosis, dysregulated vascularization including ischemic coronary artery disease, myocardioinfarction, ischemia, pulmonary hypertension, stroke, emphysema, anemia, and restenosis following angioplasty or vascular surgery;
- [035] an immune and/or inflammatory disorder, especially those involving immunological defects associated with aberrant B & T cell development. Such disorders can include, but are not limited to, acquired immunodeficiency syndrome (AIDS), Addison's disease, adult respiratory distress syndrome, allergies, ankylosing spondylitis, amyloidosis, asthma, autoimmune hemolytic anemia, autoimmune thyroiditis, Crohn's disease, episodic lymphopenia with lymphocytotoxins, erythroblastosis fetalis, Goodpasture's syndrome, Graves' disease, Hashimoto's thyroiditis, hypereosinophilia, irritable bowel syndrome and other interbowel diseases, Lupus, myasthenia gravis, myocardial or pericardial inflammation, pancreatitis, polymyositis, psoriasis, Reiter's syndrome, scleroderma, systemic anaphylaxis, ulcerative colitis, nephritis (including glomerulonephritis), gout, arthritis (such as rheumatoid arthritis and osteoarthritis), erythema, dermatitis, dermatomyositis, bronchitis, cholecystitis, and gastritis;
- [036] a disorder of carbohydrate metabolism, including but not limited to diabetic (including diabetes mellitus, diabetic retinopathy);
- [037] a parasitic, fungal and/or viral infection, including but not limited to herpes virus, poxvirus, Epstein-Barr virus, Sindbis virus and adenovirus, HIV, human papilloma virus; and
- [038] a neurodegenerative disorder, including but not limited to, Alzheimer's disease, amyotrophic lateral sclerosis, retinitis pigmentosa, Parkinson's disease, AIDS-related dementia, spinal muscular atrophy and cerebellar degeneration, myelodysplastic

syndromes, degenerative diseases of the musculoskeletal system (including but not limited to osteoporosis), cystic fibrosis, macular degeneration, and multiple sclerosis.

[039] The present invention comprises compounds for modulating protein kinase activity, more particularly Checkpoint kinase activity, even more particularly Chk1 and Chk2, even still more particularly Chk1, of Formula I,



or a pharmaceutically acceptable salt, hydrate, or prodrug thereof, wherein,

R¹ is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl;

R² and R³ are each independently selected from -H, optionally substituted alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; or

R² and R³, together with the nitrogen to which they are attached, are combined to form an optionally substituted five- to seven-membered heterocyclyl, said five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

R⁴ is -H or R⁵;

R⁵ is selected from optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; or

R⁴ and R⁵, when taken together with a common nitrogen to which they are attached, form an optionally substituted five- to seven-membered heterocyclyl, said optionally substituted five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

Z is -N= or -C(H)=;

A is selected from $-C(=O)-$, $-C(=S)-$, $-C(=NR^6)-$, and $-R^7$, when A is $-R^7$, E does not exist;

R^6 is selected from $-H$, $-NO_2$, $-N(R^2)R^3$, $-NC(=NR^2)N(R^2)R^3$, $-CN$, $-OR^4$, optionally substituted lower alkyl, optionally substituted heteroalicycylalkyl, optionally substituted aryl, optionally substituted arylalkyl and optionally substituted heteroalicyclic;

R^7 is an optionally substituted five- to seven-membered heterocyclyl, said five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

E is selected from $-N(R^8)R^9$, $-NN(R^2)R^3$, $-NNC(=O)N(R^2)R^3$, $-OR^4$, and $-R^{10}$;

R^8 is $-H$ or optionally substituted alkyl;

R^9 is selected from $-H$, optionally substituted heteroalicycylalkyl, optionally substituted heteroarylalkyl, and optionally substituted heteroalicyclic; or

R^8 and R^9 , together with the nitrogen to which they are attached, are combined to form an optionally substituted five- to seven-membered heteroalicycyl, said five- to seven-membered heteroalicycyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

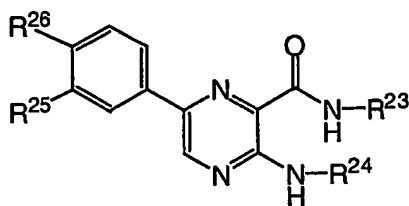
R^{10} is selected from $-H$, optionally substituted heteroalicycylalkyl, optionally substituted heteroalicyclic, optionally substituted arylalkyl, optionally substituted aryl, and optionally substituted alkyl;

W is either an optionally substituted six- to ten-membered arylene or an optionally substituted five- to ten-membered heteroarylene;

X is selected from a single bond, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}CO_2(CH_2)_{0-3}-$, $-(CH_2)_{0-3}SO_2N(R^4)(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}N(R^4)C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}S(O)_{0-2}(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{1-3}C(=O)-$, $-(CH_2)_{0-3}OC(=O)N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)N(R^4)-$, $-(CH_2)_{0-3}CO_2(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}O-$, $-(CH_2)_{0-3}N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)(CH_2)_{2-3}N(R^4)-$, $-O(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}S(O)_{0-2}-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}S(O)_{0-2}-$, $-(CH_2)_{0-3}O(CH_2)_{0-3}-$, optionally substituted alkoxy, and optionally substituted lower alkylene, wherein any $-CH_2-$ in X is optionally substituted; and

Y is selected from $-H$, optionally substituted lower alkyl, optionally substituted aryl, and optionally substituted heterocyclyl;

provided the compound is not:



wherein R^{23} , R^{24} , and R^{25} are -H; and R^{26} is -H, -Cl, -CH₃, or -OCH₃; or

wherein R^{23} is -C(NH)NH₂; R^{24} is -CH₃; R^{25} is H; and R^{26} is -Cl; or

wherein R^{23} is -H, methyl, ethyl, propyl, isopropyl, butyl, t-butyl, 1-methylpropyl, hydroxyethyl; and R^{24} , R^{25} , and R^{26} are H; or

wherein R^{23} is -H; R^{24} is butyl; and R^{25} and R^{26} are methoxy; or

wherein R^{23} 1-methylpropyl; R^{24} is -CHO; R^{25} is -H; and R^{26} is -Cl; or

wherein R^{23} is -H; R^{24} is -CH₃; and R^{25} and R^{26} are -H.

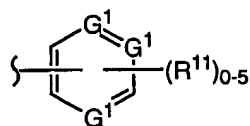
[040] In one example, the compound is according to paragraph [0039], wherein Z is =N-.

[041] In another example, the compound is according to paragraph [0040], wherein R^1 is -H.

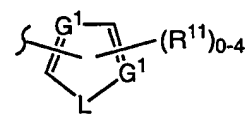
[042] In another example, the compound is according to paragraph [0041], wherein R^2 is -H and R^3 is either -H or lower alkyl.

[043] In another example, the compound is according to paragraph [0042], wherein R^2 and R^3 are -H.

[044] In another example, the compound is according to paragraph [0043], wherein A is R^7 , and R^7 is either



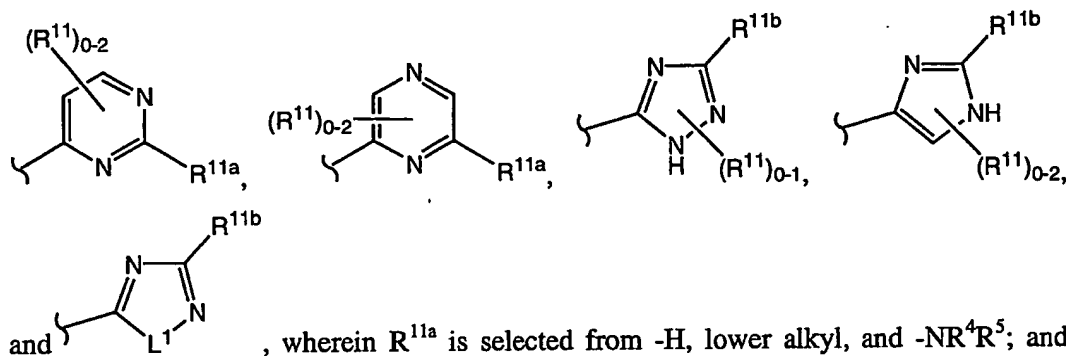
or



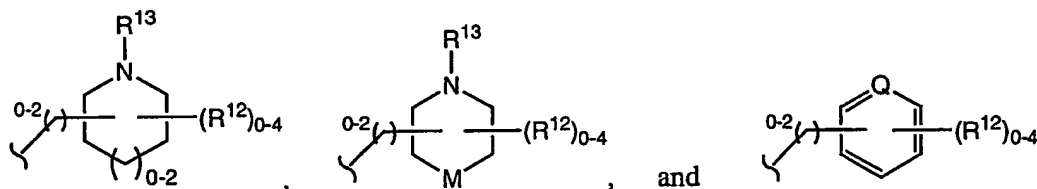
wherein each G^1 is independently =N- or =C(R^{11})-, L is selected from -O-, -S(O)₀₋₂-, and -NR¹¹, wherein each R^{11} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R^4)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R^4)R⁵, -CO₂R⁴, -C(O)N(R^4)R⁵, -N(R^4)SO₂R⁵, -N(R^4)C(O)R⁵, -N(R^4)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; or optionally two R^{11} 's

form a ring system fused with the existing ring of R^7 , said ring system substituted with 0 to 3 additional of R^{11} and said ring system optionally containing between 1 and 3 heteroatoms.

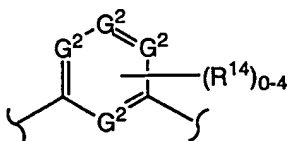
[045] In another example, the compound is according to paragraph [0044], wherein R^7 is selected from



[046] In another example, the compound is according to paragraph [0045], wherein R^{11b} is selected from optionally substituted alkyl,

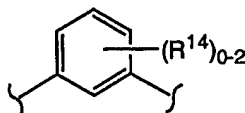


[047] In another example, the compound is according to paragraph [0046], wherein W is



wherein each G^2 is independently $=N-$ or $=C(R^{14})-$, wherein each R^{14} is independently selected from $-H$, halogen, $-CN$, $-NH_2$, $-CF_3$, $-NO_2$, $-OR^4$, $-N(R^4)R^5$, $-S(O)_{0-2}R^5$, $-SO_2N(R^4)R^5$, $-CO_2R^4$, $-C(O)N(R^4)R^5$, $-N(R^4)SO_2R^5$, $-N(R^4)C(O)R^5$, $-N(R^4)CO_2R^5$, $-C(O)R^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

[048] In another example, the compound is according to paragraph [0047], wherein W is



[049] In another example, the compound is according to paragraph [0048], wherein X is selected from $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}SO_2N(R^4)(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}N(R^4)C(=O)N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)N(R^4)-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}O-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{1-3}C(=O)-$, $-O(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}OC(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}O(CH_2)_{0-3}-$, $-(CH_2)_{0-3}CO_2(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}O-$, $-C(=O)N(R^4)(CH_2)_{2-3}N(R^4)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

[050] In another example, the compound is according to paragraph [0049], wherein X is selected from $-C(=O)N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}C(=O)N(H)-$, $-C(=O)N(H)-$, $-SO_2N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}SO_2N(H)-$, $-SO_2N(H)-$, $-N(H)C(=O)N(H)(CH_2)_{1-2}-$, $-N(H)C(=O)N(H)-$, $-(CH_2)_{0-2}N(R^4)(CH_2)_{0-2}-$, $-(CH_2)_{0-2}O(CH_2)_{0-2}-$, $-C(=O)N(H)N(H)-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-OCH_2C(=O)N(H)CH_2-$, $-(CH_2)_{1-2}N(H)C(=O)N(H)-$, $-OCH_2CH_2O-$, $-N(R^4)CH_2C(=O)N(H)CH_2-$, $-(CH_2)_{0-1}C(=O)N(H)(CH_2)_{1-2}C(=O)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

[051] In another example, the compound is according to paragraph [0050], wherein Y is selected from optionally substituted cycloalkyl, optionally substituted phenyl, and optionally substituted five- to six-membered heterocyclyl.

[052] In another example, the compound is according to paragraph [0051], wherein Y is selected from Table 1.

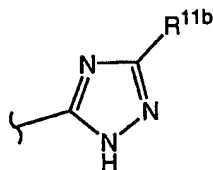
Table 1

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally

substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

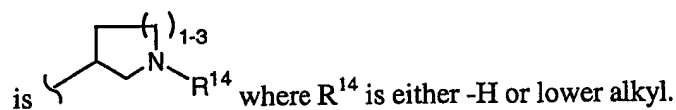
[053] In another example, the compound is according to paragraph [0052], wherein R^{11b} is selected from optionally substituted alkyl, optionally substituted phenyl, optionally substituted pyridyl, and optionally substituted heteroalicyclic.

[054] In another example, the compound is according to paragraph [0053], wherein R^7 is



[055] In another example, the compound is according to paragraph [0054], wherein R^{11b} is selected from optionally substituted morpholine, optionally substituted pyrrolidine, optionally substituted piperidine, and optionally substituted azepane.

[056] In another example, the compound is according to paragraph [0055], wherein R^{11b}



[057] In another example, the compound is according to paragraph [0054], wherein R^{11b} is a pyridine.

[058] In another example, the compound is according to paragraph [0057], wherein R^{11b} is a pyridine bearing a 2-amino group.

[059] In another example, the compound is according to paragraph [0042], wherein A is $-C(=O)-$ or $-C(=NR^6)-$.

[060] In another example, the compound is according to paragraph [0059], wherein A is $-C(=O)-$.

[061] In another example, the compound is according to paragraph [0060], wherein E is selected from Table 2.

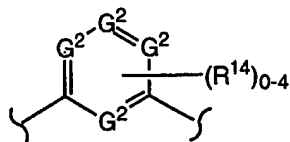
Table 2

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wherein each R^{16} is independently selected from -H, optionally substituted lower alkyl, $-\text{CO}_2R^4$, and $-\text{C}(=\text{O})R^4$; or optionally two of R^{16} , together with the nitrogen or nitrogens to which they are attached, form a heterocyclic ring; each R^{17} is independently selected from -H, halogen, oxo, -CN, $-\text{NH}_2$, $-\text{CF}_3$, $-\text{NO}_2$, $-\text{OR}^4$, $-\text{N}(\text{R}^4)\text{R}^5$, $-\text{S}(\text{O})_{0-2}\text{R}^5$, $-\text{SO}_2\text{N}(\text{R}^4)\text{R}^5$, $-\text{CO}_2\text{R}^4$, $-\text{C}(\text{O})\text{N}(\text{R}^4)\text{R}^5$, $-\text{N}(\text{R}^4)\text{SO}_2\text{R}^5$, $-\text{N}(\text{R}^4)\text{C}(\text{O})\text{R}^5$, $-\text{N}(\text{R}^4)\text{CO}_2\text{R}^5$, $-\text{C}(\text{O})\text{R}^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; R^{18} is optionally substituted lower alkyl; J^1 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0]; and J^2 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected

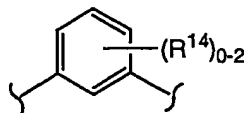
from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0], wherein the nitrogen bearing R^{16} and any nitrogen in J^2 must have at least two carbons between them.

[062] In another example, the compound is according to paragraph [0061], wherein W is



wherein each G^2 is independently $=N-$ or $=C(R^{14})-$, wherein each R^{14} is independently selected from $-H$, halogen, $-CN$, $-NH_2$, $-CF_3$, $-NO_2$, $-OR^4$, $-N(R^4)R^5$, $-S(O)_{0-2}R^5$, $-SO_2N(R^4)R^5$, $-CO_2R^4$, $-C(O)N(R^4)R^5$, $-N(R^4)SO_2R^5$, $-N(R^4)C(O)R^5$, $-N(R^4)CO_2R^5$, $-C(O)R^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

[063] In another example, the compound is according to paragraph [0062], wherein W is



[064] In another example, the compound is according to paragraph [0063], wherein X is selected from $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}SO_2N(R^4)(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}N(R^4)C(=O)N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)N(R^4)-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}O-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{1-3}C(=O)-$, $-O(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}OC(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}O(CH_2)_{0-3}-$, $-(CH_2)_{0-3}CO_2(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}O-$, $-C(=O)N(R^4)(CH_2)_{2-3}N(R^4)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

[065] In another example, the compound is according to paragraph [0064], wherein X is selected from $-C(=O)N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}C(=O)N(H)-$, $-C(=O)N(H)-$, $-SO_2N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}SO_2N(H)-$, $-SO_2N(H)-$, $-N(H)C(=O)N(H)(CH_2)_{1-2}-$, $-N(H)C(=O)N(H)-$, $-(CH_2)_{0-2}N(R^4)(CH_2)_{0-2}-$, $-(CH_2)_{0-2}O(CH_2)_{0-2}-$, $-C(=O)N(H)N(H)-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-OCH_2C(=O)N(H)CH_2-$, $-(CH_2)_{1-2}N(H)C(=O)N(H)-$, $-OCH_2CH_2O-$, $-N(R^4)CH_2C(=O)N(H)CH_2-$, $-(CH_2)_{0-1}C(=O)N(H)(CH_2)_{1-2}C(=O)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

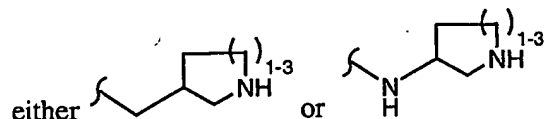
[066] In another example, the compound is according to paragraph [0065], wherein Y is selected from optionally substituted cycloalkyl, optionally substituted phenyl, and optionally substituted five- to six-membered heterocyclyl.

[067] In another example, the compound is according to paragraph [0066], wherein Y is selected from Table 3.

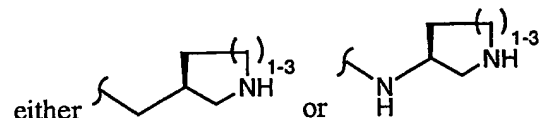
Table 3

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl.

[068] In another example, the compound is according to paragraph [0067], wherein E is



[069] In another example, the compound is according to paragraph [0068], wherein E is

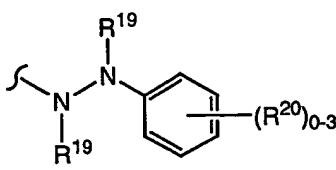
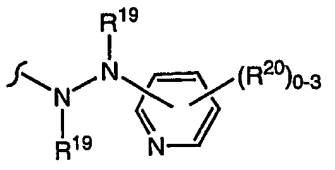
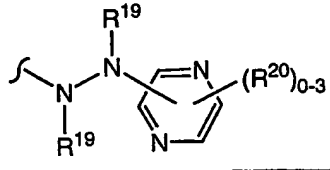
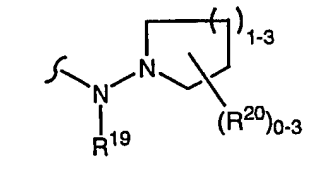
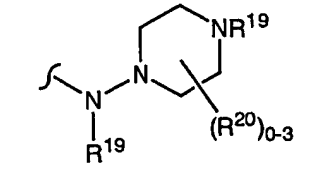
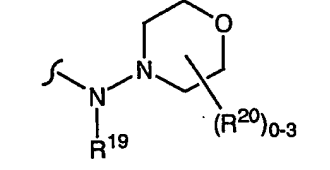
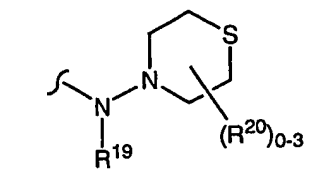
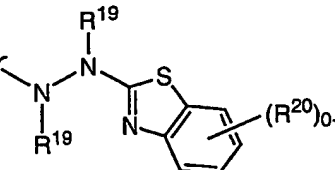
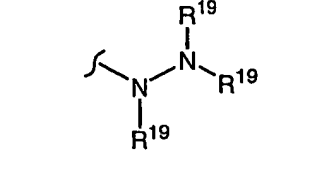


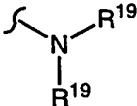
[070] In another example, the compound is according to paragraph [0069], wherein R² and R³ are -H.

[071] In another example, the compound is according to paragraph [0070], wherein A is -C(=NR⁶)-.

[072] In another example, the compound is according to paragraph [0071], wherein E is selected from -H, lower alkyl, and a group listed in Table 4,

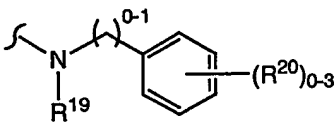
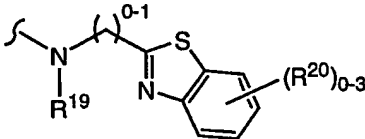
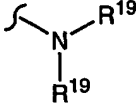
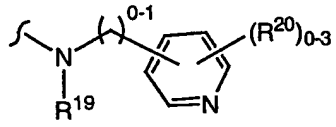
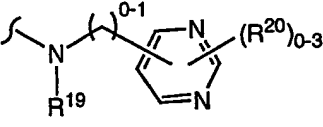
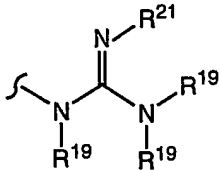
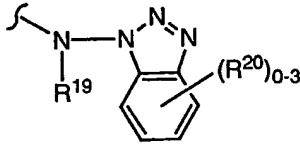
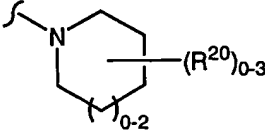
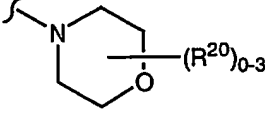
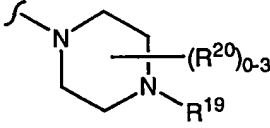
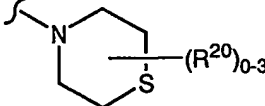
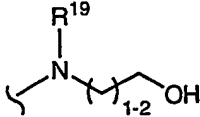
Table 4

		
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R^6 is selected from -H, -OH, lower alkyl, -NO₂, -CN, lower alkoxy, and a group listed in Table 5,

Table 5

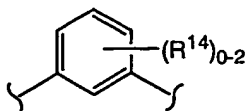
		
		
		
		

wherein each R^{19} is independently either -H or lower alkyl;

each R^{20} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl;

and R^{21} is selected from -H, -OH, lower alkyl, -NO₂, -CN, lower alkoxy.

[073] In another example, the compound is according to paragraph [0072], wherein W is



[074] In another example, the compound is according to paragraph [0073], wherein X is selected from $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}SO_2N(R^4)(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}N(R^4)C(=O)N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)N(R^4)-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}O-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{1-3}C(=O)-$, $-O(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}OC(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}O(CH_2)_{0-3}-$, $-(CH_2)_{0-3}CO_2(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}O-$, $-C(=O)N(R^4)(CH_2)_{2-3}N(R^4)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

[075] In another example, the compound is according to paragraph [0074], wherein X is selected from $-C(=O)N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}C(=O)N(H)-$, $-C(=O)N(H)-$, $-SO_2N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}SO_2N(H)-$, $-SO_2N(H)-$, $-N(H)C(=O)N(H)(CH_2)_{1-2}-$, $-N(H)C(=O)N(H)-$, $-(CH_2)_{0-2}N(R^4)(CH_2)_{0-2}-$, $-(CH_2)_{0-2}O(CH_2)_{0-2}-$, $-C(=O)N(H)N(H)-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-OCH_2C(=O)N(H)CH_2-$, $-(CH_2)_{1-2}N(H)C(=O)N(H)-$, $-OCH_2CH_2O-$, $-N(R^4)CH_2C(=O)N(H)CH_2-$, $-(CH_2)_{0-1}C(=O)N(H)(CH_2)_{1-2}C(=O)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

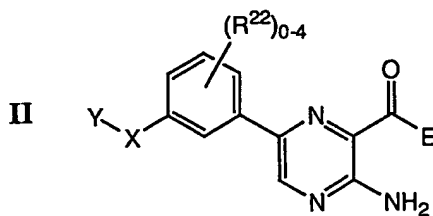
[076] In another example, the compound is according to paragraph [0075], wherein Y is selected from optionally substituted cycloalkyl, optionally substituted phenyl, and optionally substituted five- to six-membered heterocyclyl.

[077] In another example, the compound is according to paragraph [0076], wherein Y is selected from Table 6.

Table 6

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

[078] In another example, the compound is according to paragraph [0039], of formula II,



wherein E is selected from Table 7,

Table 7

wherein each R^{16} is independently selected from -H, optionally substituted lower alkyl, $-\text{CO}_2R^4$, and $-\text{C}(=\text{O})R^4$;

or optionally two of R^{16} , together with the nitrogen or nitrogens to which they are attached, form a heterocyclic ring;

each R^{17} is independently selected from -H, halogen, oxo, -CN, $-\text{NH}_2$, $-\text{CF}_3$, $-\text{NO}_2$, $-\text{OR}^4$, $-\text{N}(\text{R}^4)\text{R}^5$, $-\text{S}(\text{O})_{0.2}\text{R}^5$, $-\text{SO}_2\text{N}(\text{R}^4)\text{R}^5$, $-\text{CO}_2R^4$, $-\text{C}(\text{O})\text{N}(\text{R}^4)\text{R}^5$, $-\text{N}(\text{R}^4)\text{SO}_2\text{R}^5$, $-\text{N}(\text{R}^4)\text{C}(\text{O})\text{R}^5$, $-\text{N}(\text{R}^4)\text{CO}_2\text{R}^5$, $-\text{C}(\text{O})\text{R}^4$, optionally substituted lower alkyl, optionally substituted aryl,

optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl;

R^{18} is optionally substituted lower alkyl;

J^1 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0];

J^2 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0], wherein the nitrogen bearing R^{16} and any nitrogen in J^2 must have at least two carbons between them.

X is selected from $-C(=O)N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}C(=O)N(H)-$, $-C(=O)N(H)-$, $-SO_2N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}SO_2N(H)-$, $-SO_2N(H)-$, $-N(H)C(=O)N(H)(CH_2)_{1-2}-$, $-N(H)C(=O)N(H)-$, $-(CH_2)_{0-2}N(R^4)(CH_2)_{0-2}-$, $-(CH_2)_{0-2}O(CH_2)_{0-2}-$, $-C(=O)N(H)N(H)-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-OCH_2C(=O)N(H)CH_2-$, $-(CH_2)_{1-2}N(H)C(=O)N(H)-$, $-OCH_2CH_2O-$, $-N(R^4)CH_2C(=O)N(H)CH_2-$, $-(CH_2)_{0-1}C(=O)N(H)(CH_2)_{1-2}C(=O)-$, optionally substituted alkoxy, and optionally substituted lower alkylene;

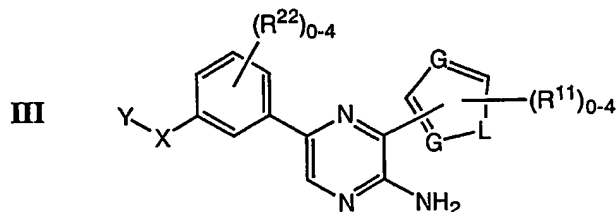
Y is selected from Table 8,

Table 8

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl; and,

each R^{22} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

[079] In another example, the compound is according to paragraph [0039], of formula III,



wherein, each G is independently =N- or =C(R¹¹)-;

L is selected from -O-, -S(O)₀₋₂-, and -NR¹¹;

each R^{11} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; or

optionally two R^{11} 's form a ring system fused with the existing ring of R^7 , said ring system substituted with 0 to 3 additional of R^{11} and said ring system optionally containing between 1 and 3 heteroatoms;

X is selected from -C(=O)N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂C(=O)N(H)-, -C(=O)N(H)-, -SO₂N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂SO₂N(H)-, -SO₂N(H)-, -N(H)C(=O)N(H)(CH₂)₁₋₂-, -N(H)C(=O)N(H)-, -(CH₂)₀₋₂N(R⁴)(CH₂)₀₋₂-, -(CH₂)₀₋₂O(CH₂)₀₋₂-, -C(=O)N(H)N(H)-, -N(R⁴)(CH₂)₂₋₃N(R⁴)-, -OCH₂C(=O)N(H)CH₂-, -(CH₂)₁₋₂N(H)C(=O)N(H)-, -OCH₂CH₂O-, -N(R⁴)CH₂C(=O)N(H)CH₂-, -(CH₂)₀₋₁C(=O)N(H)(CH₂)₁₋₂C(=O)-, optionally substituted alkoxy, and optionally substituted lower alkylene;

Y is selected from Table 9,

Table 9

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; and,

each R^{22} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl.

[080] In another example, the compound is according to paragraph [0039], wherein the compound is selected from Table 10.

Table 10

#	Name
1	3-amino-6-phenyl-N-(2-phenylethyl)pyrazine-2-carboxamide
2	3-amino-6-phenyl-N-(phenylmethyl)pyrazine-2-carboxamide
3	3-amino-6-(3-chlorophenyl)pyrazine-2-carboxamide
4	3-amino-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide
5	3-amino-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide
6	3-amino-N-(2,3-dihydro-1H-inden-2-yl)-6-phenylpyrazine-2-carboxamide

#	Name
7	3-amino-6-(2-chlorophenyl)pyrazine-2-carboxamide
8	3-amino-N-cyclopentyl-6-phenylpyrazine-2-carboxamide
9	3-amino-N,N-dimethyl-6-phenylpyrazine-2-carboxamide
10	3-amino-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide
11	5-phenyl-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine
12	3-amino-N-(cyclopropylmethyl)-6-phenylpyrazine-2-carboxamide
13	3-amino-6-phenyl-N-(tetrahydrofuran-2-ylmethyl)pyrazine-2-carboxamide
14	3-amino-N-cyclohexyl-6-phenylpyrazine-2-carboxamide
15	6-phenyl-N-propyl-3-(propylamino)pyrazine-2-carboxamide
16	3-amino-6-(2-methylphenyl)pyrazine-2-carboxamide
17	3-amino-6-[2-(methyloxy)phenyl]pyrazine-2-carboxamide
18	3-amino-6-(2,4-difluorophenyl)pyrazine-2-carboxamide
19	3-amino-6-(3-fluorophenyl)pyrazine-2-carboxamide
20	3-amino-N-cyclopropyl-6-phenylpyrazine-2-carboxamide
21	3-amino-6-(3,4-difluorophenyl)pyrazine-2-carboxamide
22	3-amino-6-(4-fluorophenyl)-N-methylpyrazine-2-carboxamide
23	3-amino-6-[4-(ethyloxy)phenyl]pyrazine-2-carboxamide
24	3-amino-6-[3-(ethyloxy)phenyl]pyrazine-2-carboxamide
25	3-amino-6-(1,3-benzodioxol-5-yl)pyrazine-2-carboxamide
26	3-amino-6-naphthalen-1-ylpyrazine-2-carboxamide
27	3-amino-6-naphthalen-2-ylpyrazine-2-carboxamide
28	3-amino-6-[3-[(trifluoromethyl)oxy]phenyl]pyrazine-2-carboxamide
29	3-amino-6-[3-(aminocarbonyl)phenyl]pyrazine-2-carboxamide

#	Name
30	6-[3-(acetylamino)phenyl]-3-aminopyrazine-2-carboxamide
31	3-amino-6-biphenyl-4-ylpyrazine-2-carboxamide
32	3-amino-6-[4-(dimethylamino)phenyl]pyrazine-2-carboxamide
33	3-amino-6-(3-methylphenyl)pyrazine-2-carboxamide
34	methyl 3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoate
35	3-amino-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide
36	3-amino-6-(3-hydroxyphenyl)pyrazine-2-carboxamide
37	3-amino-6-(1-benzofuran-2-yl)pyrazine-2-carboxamide
38	3-(methylamino)-6-phenylpyrazine-2-carboxamide
39	6-phenyl-3-[(phenylmethyl)amino]pyrazine-2-carboxamide
40	6-phenyl-3-(propylamino)pyrazine-2-carboxamide
41	3-amino-6-biphenyl-3-ylpyrazine-2-carboxamide
42	3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoic acid
43	3-amino-6-{4-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide
44	3-amino-N-methyl-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide
45	3-amino-6-(3-hydroxyphenyl)-N-methylpyrazine-2-carboxamide
46	3-amino-N-[2-(methyloxy)ethyl]-6-phenylpyrazine-2-carboxamide
47	N-[2-(acetylamino)ethyl]-3-amino-6-phenylpyrazine-2-carboxamide
48	3-amino-6-phenylpyrazine-2-carbohydrazide
49	3-amino-N-hydroxy-6-phenylpyrazine-2-carboxamide
50	3-amino-6-[3-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide
51	3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide
52	3-amino-6-[4-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide

#	Name
53	3-amino-N-methyl-6-(3-methylphenyl)pyrazine-2-carboxamide
54	3-amino-N-methyl-6-[4-(methyloxy)phenyl]pyrazine-2-carboxamide
55	3-amino-6-biphenyl-3-yl-N-methylpyrazine-2-carboxamide
56	6-[3-(acetylamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide
57	3-amino-6-[3-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide
58	3-amino-6-[4-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide
59	3-amino-N-methyl-6-[3-[(1E)-N-methylethanimidoyl]phenyl]pyrazine-2-carboxamide
60	3-amino-N-methyl-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide
61	3-amino-6-(4-fluoro-3-methylphenyl)-N-methylpyrazine-2-carboxamide
62	3-amino-N-methyl-6-pyridin-3-ylpyrazine-2-carboxamide
63	6-[4-(acetylamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide
64	3-amino-N-methyl-6-[3-[(methylamino)carbonyl]phenyl]pyrazine-2-carboxamide
65	6-(3-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide
66	3-amino-N-methyl-6-(5-pyridin-3-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
67	3-amino-6-(5-cyclopentyl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide
68	3-amino-6-(5-cyclohexyl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide
69	3-amino-N-methyl-6-(5-phenyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
70	3-amino-N-methyl-6-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide
71	3-amino-N-methyl-6-(5-methyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
72	3-amino-N-methyl-6-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide
73	3-amino-N-methyl-6-(5-pyridin-4-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
74	3-amino-6-[5-(1,1-dimethylethyl)-1,2,4-oxadiazol-3-yl]-N-methylpyrazine-2-carboxamide
75	3-amino-6-(5-furan-2-yl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide

#	Name
76	3-amino-6-[4-(dimethylamino)phenyl]-N-methylpyrazine-2-carboxamide
77	3-amino-N-methyl-6-[4-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide
78	3-amino-6-(3,5-dimethylphenyl)-N-methylpyrazine-2-carboxamide
79	6-(4-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide
80	3-amino-6-[3,4-bis(methyloxy)phenyl]-N-methylpyrazine-2-carboxamide
81	3-amino-N-methyl-6-[3-[(phenylcarbonyl)amino]phenyl]pyrazine-2-carboxamide
82	3-amino-N-methyl-6-[3-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide
83	3-amino-N-methyl-6-[3-(1H-tetrazol-5-yl)phenyl]pyrazine-2-carboxamide
84	3-amino-N-methyl-6-[3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
85	3-amino-6-[3-(aminocarbonyl)phenyl]-N-methylpyrazine-2-carboxamide
86	3-amino-6-[3-[(dimethylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
87	3-amino-N-methyl-6-[4-(methylsulfonyl)phenyl]pyrazine-2-carboxamide
88	3-amino-6-(3-aminophenyl)-N-ethylpyrazine-2-carboxamide
89	3-amino-6-[3-[[[(4-fluorophenyl)amino]carbonyl]amino]phenyl]-N-methylpyrazine-2-carboxamide
90	3-amino-N-methyl-6-[3-[[[(4-(methyloxy)phenyl)amino]carbonyl]amino]phenyl]pyrazine-2-carboxamide
91	6,6'-[(oxomethanediy)bis(iminobenzene-3,1-diyl)]bis(3-amino-N-methylpyrazine-2-carboxamide)
92	3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoic acid
93	3-amino-N-methyl-6-[3-[(4-methylpiperazin-1-yl)carbonyl]phenyl]pyrazine-2-carboxamide
94	3-amino-N-methyl-6-[3-[[[(2-phenylethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
95	3-amino-N-methyl-6-[3-[[[(4-(methyloxy)phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
96	3-amino-6-[3-[[[(1R)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
97	3-amino-6-[3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
98	3-amino-6-[3-[[[(4-chlorophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
99	3-amino-N-methyl-6-(3-([methyl(phenylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide
100	3-amino-6-[3-(((2,4-dichlorophenyl)methyl]amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
101	3-amino-N-methyl-6-[3-(((4-methylphenyl)methyl]amino)carbonyl)phenyl]pyrazine-2-carboxamide
102	3-amino-N-methyl-6-[3-(((4-(trifluoromethyl)phenyl)methyl]amino)carbonyl)phenyl]pyrazine-2-carboxamide
103	3-amino-6-[3-(((4-fluorophenyl)methyl]amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
104	3-amino-6-[3-([2,3-dihydro-1H-inden-2-ylamino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
105	3-amino-N-methyl-6-[3-([3-(methyloxy)phenyl]carbonyl)amino]phenyl]pyrazine-2-carboxamide
106	3-amino-N-methyl-6-[3-([4-(methyloxy)phenyl]carbonyl)amino]phenyl]pyrazine-2-carboxamide
107	3-amino-N-methyl-6-[3-([3-(methyloxy)phenyl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
108	3-amino-N-methyl-6-[3-([1-(4-(methyloxy)phenyl)ethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
109	3-amino-N-methyl-6-[3-([2-(trifluoromethyl)phenyl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
110	3-amino-6-[3-([3-(3-fluorophenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
111	3-amino-6-[3-([4-(4-chlorophenyl)carbonyl]amino)phenyl]-N-methylpyrazine-2-carboxamide
112	3-amino-6-[3-([1-(4-fluorophenyl)ethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
113	3-amino-6-[3-([2,3-bis(methyloxy)phenyl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
114	3-amino-6-[3-([bis(phenylmethyl)amino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
115	3-amino-6-[3-([2,4-bis(methyloxy)phenyl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
116	3-amino-6-[3-([ethyl(pyridin-4-ylmethyl)amino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
117	3-amino-N-methyl-6-[3-([4-(phenylmethyl)piperazin-1-yl]carbonyl)phenyl]pyrazine-2-carboxamide
118	3-amino-N-methyl-6-[3-([1-(phenylmethyl)piperidin-4-yl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
119	3-amino-N-methyl-6-[3-([4-(4-methylphenyl)carbonyl]amino)phenyl]pyrazine-2-carboxamide
120	3-amino-N-methyl-6-[3-([5-methylpyrazin-2-yl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
121	3-amino-N-methyl-6-[3-([pyridin-2-ylmethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide

#	Name
122	3-amino-N-methyl-6-(3-(((pyridin-3-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
123	3-amino-N-methyl-6-(3-(((pyridin-4-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
124	3-amino-6-(3-((furan-2-ylcarbonyl)amino)phenyl)-N-methylpyrazine-2-carboxamide
125	3-amino-6-(3-(((3-chlorophenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
126	3-amino-6-(3-(((cyclohexylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
127	3-amino-N-methyl-6-(3-(((2-phenylpropyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
128	3-amino-6-(3-(((2-chlorophenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
129	3-amino-6-(3-(((2-(2-chlorophenyl)ethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
130	3-amino-N-methyl-6-(3-(((2-(2-(methyloxy)phenyl)ethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
131	3-amino-6-(3-(((2-(4-fluorophenyl)ethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
132	3-amino-N-methyl-6-(3-(((2-(2-thienyl)ethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
133	3-amino-6-(3-((cyclohexylamino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
134	1,1-dimethylethyl 4-(((3-(5-amino-6-((methylamino)carbonyl)pyrazin-2-yl)phenyl)carbonyl)amino)methyl)piperidine-1-carboxylate
135	3-amino-N-methyl-6-(3-(((piperidin-4-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
136	3-amino-N-methyl-6-(3-(5-phenyl-1,2,4-oxadiazol-3-yl)phenyl)pyrazine-2-carboxamide
137	3-amino-N-methyl-6-(3-(5-(phenylmethyl)-1,2,4-oxadiazol-3-yl)phenyl)pyrazine-2-carboxamide
138	3-amino-N-methyl-6-(3-(5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl)phenyl)pyrazine-2-carboxamide
139	3-amino-6-(3-(((1,1-dimethylethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
140	3-amino-6-(3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
141	3-amino-6-(3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
142	3-amino-N-methyl-6-(3-((phenylamino)carbonyl)phenyl)pyrazine-2-carboxamide
143	3-amino-6-(3-(1,3-dihydro-2H-isindol-2-ylcarbonyl)phenyl)-N-methylpyrazine-2-carboxamide
144	3-amino-N-methyl-6-(3-(((1R)-1-phenylethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide

#	Name
145	3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
146	3-amino-N-methyl-6-[3-(((1R)-1-phenylpropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
147	3-amino-N-methyl-6-[3-(((1S)-1-phenylpropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
148	3-amino-N-methyl-6-[3-(((1R,2S)-2-phenylcyclopropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
149	phenylmethyl 4-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoate
150	3-amino-N-methyl-6-[4-(((phenylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
151	3-amino-N-methyl-6-[4-[(2-phenylethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
152	3-amino-6-[4-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
153	3-amino-N-methyl-6-[4-(((4-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
154	3-amino-6-[4-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
155	3-amino-6-[3-[(4-chlorophenyl)acetyl]amino]phenyl]-N-methylpyrazine-2-carboxamide
156	3-amino-N-methyl-6-[3-([3-(phenyloxy)phenyl]carbonyl)amino]phenyl]pyrazine-2-carboxamide
157	3-amino-6-[3-[(diphenylacetyl)amino]phenyl]-N-methylpyrazine-2-carboxamide
158	3-amino-N-methyl-6-[3-[(phenylacetyl)amino]phenyl]pyrazine-2-carboxamide
159	3-amino-N-methyl-6-[3-[(3-phenylpropanoyl)amino]phenyl]pyrazine-2-carboxamide
160	3-amino-N-methyl-6-[3-(((1R)-1,2,3,4-tetrahydronaphthalen-1-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide
161	3-amino-N-methyl-6-[3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide
162	3-amino-N-methyl-6-[3-([3,4,5-tris(methyloxy)phenyl]methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
163	3-amino-N-methyl-6-[3-(((1R,2R)-2-[(phenylmethyl)oxy]cyclopentyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
164	3-amino-N-methyl-6-[3-(((1S,2S)-2-[(phenylmethyl)oxy]cyclopentyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
165	3-amino-6-(3-cyanophenyl)-N-methylpyrazine-2-carboxamide
166	3-amino-6-[3-[(cyclopentylcarbonyl)amino]phenyl]-N-methylpyrazine-2-carboxamide
167	3-amino-6-[3-[(4-cyanophenyl)carbonyl]amino]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
168	3-amino-N-methyl-6-[3-(((phenylmethyl)amino)carbonyl)amino]phenyl]pyrazine-2-carboxamide
169	3-amino-N-methyl-6-(3-(((naphthalen-2-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
170	3-amino-N-methyl-6-(3-(((tetrahydrofuran-2-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
171	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
172	3-amino-6-[3-(((3-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
173	3-amino-6-[3-(((2-[3,5-bis(methyloxy)phenyl]ethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
174	3-amino-N-methyl-6-[3-((2-phenylethyl)oxy)phenyl]pyrazine-2-carboxamide
175	3-amino-N-methyl-6-[3-((2-(4-methylpiperidin-1-yl)ethyl)oxy)phenyl]pyrazine-2-carboxamide
176	3-amino-6-[3-((2-[4-(2-hydroxyethyl)piperidin-1-yl]ethyl)oxy)phenyl]-N-methylpyrazine-2-carboxamide
177	3-amino-6-[3-((2-(3-hydroxypyrrolidin-1-yl)ethyl)oxy)phenyl]-N-methylpyrazine-2-carboxamide
178	3-amino-6-[3-((2-[ethyl(phenylmethyl)amino]ethyl)oxy)phenyl]-N-methylpyrazine-2-carboxamide
179	3-amino-6-[3-((2-(diethylamino)ethyl)oxy)phenyl]-N-methylpyrazine-2-carboxamide
180	3-amino-6-[3-(((biphenyl-2-ylmethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
181	3-amino-N-methyl-6-[3-((2-(4-phenylpiperazin-1-yl)ethyl)oxy)phenyl]pyrazine-2-carboxamide
182	3-amino-6-[3-(((biphenyl-4-ylmethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
183	3-amino-N-methyl-6-[3-((2-[methyl(phenylmethyl)amino]ethyl)oxy)phenyl]pyrazine-2-carboxamide
184	3-amino-N-methyl-6-[3-(((4-(phenyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
185	3-amino-N-methyl-6-[3-((2-morpholin-4-ylethyl)oxy)phenyl]pyrazine-2-carboxamide
186	3-amino-6-[3-((2-[(cyclopropylmethyl)(propyl)amino]ethyl)oxy)phenyl]-N-methylpyrazine-2-carboxamide
187	3-amino-N-methyl-6-[3-((2-[4-(phenylmethyl)piperidin-1-yl]ethyl)oxy)phenyl]pyrazine-2-carboxamide
188	3-amino-N-methyl-6-[3-((2-piperidin-1-ylethyl)oxy)phenyl]pyrazine-2-carboxamide
189	3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
190	3-amino-6-[3-((2-(4-ethylpiperazin-1-yl)ethyl)oxy)phenyl]-N-methylpyrazine-2-carboxamide

#	Name
191	3-amino-N-methyl-6-[3-[(2-pyrrolidin-1-ylethyl)oxy]phenyl]pyrazine-2-carboxamide
192	3-amino-N-methyl-6-[3-[[2-(4-methylpiperazin-1-yl)ethyl]oxy]phenyl]pyrazine-2-carboxamide
193	3-amino-6-[3-[(2-[4-(2-hydroxyethyl)piperazin-1-yl]ethyl)oxy]phenyl]-N-methylpyrazine-2-carboxamide
194	3-amino-6-[3-[[2-(3,4-dihydroisoquinolin-2(1H)-yl)ethyl]oxy]phenyl]-N-methylpyrazine-2-carboxamide
195	ethyl 4-[2-[(3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)phenyl]oxy]ethyl]piperazine-1-carboxylate
196	3-amino-N-methyl-6-[3-(((3S)-1-(phenylmethyl)pyrrolidin-3-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
197	3-amino-N-methyl-6-[3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
198	3-amino-N-methyl-6-[3-(((2-(phenyloxy)ethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
199	3-amino-N-methyl-6-[3-(((2-[4-(phenylmethyl)piperazin-1-yl]ethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
200	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)piperidin-4-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
201	3-amino-6-[3-(((furan-3-ylmethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
202	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-2-yl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
203	3-amino-6-[3-(((furan-2-ylmethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
204	3-amino-N-methyl-6-[3-(((4-(phenylmethyl)morpholin-2-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
205	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
206	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
207	3-amino-6-[3-(((2,3-dihydro-1-benzofuran-5-yl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
208	3-amino-6-[3-(((6-fluoro-4H-1,3-benzodioxin-8-yl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
209	3-amino-6-[3-(((2,4-dichlorophenyl)acetyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
210	3-amino-6-[3-(((3,4-dichlorophenyl)acetyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
211	3-amino-N-methyl-6-[3-(((4-(trifluoromethyl)phenyl)acetyl)amino)phenyl]pyrazine-2-carboxamide
212	3-amino-6-[3-(((3,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
213	3-amino-6-[3-(((5-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
214	3-amino-6-[3-(((2,4-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
215	3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
216	3-amino-N-methyl-6-[3-(((4-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
217	3-amino-N-methyl-6-[3-(((2-methylpropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
218	3-amino-N-methyl-6-[3-(((3-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
219	3-amino-6-[3-(((cyclopropylmethyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
220	3-amino-6-[3-(((cyclopentylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
221	3-amino-N-methyl-6-[3-(((4-methylphenyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
222	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
223	3-amino-6-[3-(((1,3-benzodioxol-5-ylmethyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
224	3-amino-6-[3-(((biphenyl-3-ylmethyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
225	3-amino-N-methyl-6-[3-(((1-methyl-1-phenylethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
226	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
227	3-amino-N-methyl-6-[3-(((4-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
228	3-amino-6-[3-(((4-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
229	3-amino-6-[3-(((4-chloro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
230	3-amino-6-[3-(((4-aminophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
231	3-amino-6-[3-(((2-aminophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
232	3-amino-6-[3-(((3-aminophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
233	3-amino-N-methyl-6-pyrimidin-5-ylpyrazine-2-carboxamide
234	3-amino-N-methyl-6-[3-(methylsulfonyl)phenyl]pyrazine-2-carboxamide
235	3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide

#	Name
236	3-amino-6-(4-hydroxyphenyl)-N-methylpyrazine-2-carboxamide
237	3-amino-6-{3-[[[4-(aminosulfonyl)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
238	3-amino-6-{3-[[[4-(dimethylamino)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
239	3-amino-6-{3-[[[3-bromophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
240	3-amino-6-(4-chloro-3-[(phenylmethyl)amino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
241	methyl 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoate
242	methyl 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoate
243	3-amino-6-{3-[[[2,3-dimethylphenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
244	3-amino-6-{3-[[[4-hydroxy-3-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
245	3-amino-6-{3-[[[4-chloro-2-methylphenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
246	3-amino-6-{3-[[[2-(ethyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
247	3-amino-6-{3-[[[2-chloro-6-(phenyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
248	3-amino-6-{3-[[[2,5-dichlorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
249	3-amino-6-{3-[[[2,6-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
250	3-amino-6-{3-[[[2-chloro-6-methylphenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
251	3-amino-N-methyl-6-{3-[[[2,4,5-trifluorophenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
252	3-amino-6-{3-[[[2,2-dimethylpropyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
253	3-amino-6-{3-[[[cyclopentylmethyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
254	3-amino-6-{3-[[[3,5-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
255	3-amino-N-methyl-6-{3-[[[4-(methylsulfonyl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
256	3-amino-6-{3-[[[5-bromo-2-fluorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
257	3-amino-6-{3-[[[3-bromo-4-fluorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide
258	3-amino-N-methyl-6-{3-[[[2,4,6-tris(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide

#	Name
259	3-amino-N-methyl-6-[3-(((3-methylphenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
260	3-amino-6-[3-(((4-bromo-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
261	3-amino-6-[3-(((2,5-dimethylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
262	3-amino-6-[3-(((3,5-bis(trifluoromethyl)phenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
263	3-amino-6-[3-(((2,2-diphenylethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
264	3-amino-N-methyl-6-(3-thienyl)pyrazine-2-carboxamide
265	3-amino-N-methyl-6-[3-(((phenylmethyl)amino)sulfonyl)phenyl]pyrazine-2-carboxamide
266	5,5'-diamino-N,N'-dimethyl-2,2'-bipyrazine-6,6'-dicarboxamide
267	3-amino-6-(1H-indol-5-yl)-N-methylpyrazine-2-carboxamide
268	3-amino-N-methyl-6-(2-thienyl)pyrazine-2-carboxamide
269	3-amino-N-methyl-6-pyridin-4-ylpyrazine-2-carboxamide
270	3-amino-N-cyclopropyl-6-[3-(((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
271	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-cyclopropylpyrazine-2-carboxamide
272	3-amino-N-ethyl-6-[3-(((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
273	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-ethylpyrazine-2-carboxamide
274	3-amino-6-[3-(((2,3-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
275	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
276	3-amino-6-[3-(((3,5-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
277	3-amino-6-[3-(((5-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
278	3-amino-6-[3-((cyclopropylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
279	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
280	3-amino-6-[3-(((4'-fluorobiphenyl-2-yl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
281	3-amino-N-methyl-6-[3-(((2,3,4-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide

#	Name
282	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
283	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
284	3-amino-6-[3-((9H-fluoren-9-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
285	3-amino-N-methyl-6-[3-(((2-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
286	3-amino-6-[3-(((3,4-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
287	3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
288	3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
289	3-amino-6-[3-(((2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
290	3-amino-N-methyl-6-[3-(((4-(4-methylpiperazin-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
291	3-amino-6-(4-fluoro-3-((phenylmethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
292	3-amino-N-methyl-6-[3-(((2,4,6-trichlorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
293	3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
294	3-amino-6-[3-(((2,5-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
295	3-amino-6-[3-(((3-fluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
296	3-amino-6-[3-(((2-chloro-6-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
297	3-amino-6-[3-(((2-chloro-3,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
298	3-amino-6-[3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
299	3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
300	3-amino-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
301	3-amino-6-[3-(((6-chloro-2-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
302	3-amino-6-[3-(((4-iodophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
303	3-amino-6-[3-(((3-chloro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
304	3-amino-6-[3-(((4-bromophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
305	3-amino-6-[3-(((2-bromophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
306	3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
307	3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
308	3-amino-N-methyl-6-(3-(((phenylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
309	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
310	3-amino-6-[3-(((2,5-difluorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
311	3-amino-N-methyl-6-(3-(((naphthalen-2-ylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
312	3-amino-N-methyl-6-(3-(((phenylmethyl)thio)methyl)phenyl)pyrazine-2-carboxamide
313	3-amino-N-methyl-6-(3-(((phenylmethyl)sulfonyl)methyl)phenyl)pyrazine-2-carboxamide
314	3-amino-N-methyl-6-(3-(((phenylmethyl)oxy)methyl)phenyl)pyrazine-2-carboxamide
315	3-amino-N-methyl-6-(3-(((phenylmethyl)amino)methyl)phenyl)pyrazine-2-carboxamide
316	3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
317	9H-fluoren-9-ylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
318	phenylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
319	ethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
320	3-amino-6-[3-(((3,4-dichlorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
321	1,1-dimethylethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
322	(1R,2S,5R)-5-methyl-2-(1-methylethyl)cyclohexyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
323	3-amino-6-[3-(((4-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
324	3-amino-N-methyl-6-(3-(((4-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
325	3-amino-N-methyl-6-[3-(((2-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
326	3-amino-6-[3-(((3-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
327	3-amino-6-[3-(((2-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
328	3-amino-N-methyl-6-[3-[(2-phenylhydrazino)carbonyl]phenyl]pyrazine-2-carboxamide
329	3-amino-N-(2-hydroxyethyl)-6-(3-methylphenyl)pyrazine-2-carboxamide
330	3-amino-N-methyl-6-[3-[(phenylacetyl)amino]methyl]phenyl]pyrazine-2-carboxamide
331	3-amino-6-[3-[[[(4-chlorophenyl)acetyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
332	3-amino-6-[3-[[[(biphenyl-4-ylcarbonyl)amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
333	3-amino-6-[3-[[[(2-chloro-5-(trifluoromethyl)phenyl)carbonyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
334	N-[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)methyl]-1H-indole-2-carboxamide
335	3-amino-6-[3-[[[(3-hydroxypyridin-2-yl)carbonyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
336	3-amino-6-[3-[[[(1H-indol-3-ylacetyl)amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
337	3-amino-6-[3-[[[(2,5-difluorophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
338	3-amino-6-[3-[[[(2,6-difluorophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
339	3-amino-6-[3-[[[(3,5-difluorophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
340	3-amino-6-[3-[[[(3-iodophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
341	3-amino-N-methyl-6-[3-[[[(2-morpholin-4-ylphenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
342	3-amino-N-methyl-6-[3-[[[(3-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
343	3-amino-N-methyl-6-[3-[[[(4-morpholin-4-ylphenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
344	3-amino-N-methyl-6-[3-[[[(3-morpholin-4-ylphenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
345	3-amino-N-methyl-6-[3-[[[(4-(1-methylethyl)phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
346	3-amino-N-methyl-6-[3-[[[(4-(1-methylethyl)phenyl)oxy]acetyl]amino]methyl]phenyl]pyrazine-2-carboxamide
347	3-amino-6-[3-[[[(4-aminophenyl)thio]acetyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
348	3-amino-N-methyl-6-(4-(methyloxy)-3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
349	3-amino-N-methyl-6-(5-[[[(phenylmethyl)amino]carbonyl]pyridin-3-yl]pyrazine-2-carboxamide
350	3-amino-6-[3-[[[(4-iodophenyl)carbonyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
351	3-amino-N-methyl-6-[3-(((4-pentylphenyl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide
352	3-amino-6-(3-(((1,3-benzodioxol-5-ylcarbonyl)amino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
353	3-amino-6-(3-(((4-(1,1-dimethylethyl)phenyl)methyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
354	3-amino-N-methyl-6-(3-(((1-[5-(trifluoromethyl)pyrimidin-2-yl]piperidin-4-yl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide
355	3-amino-N-methyl-6-[3-(((2-piperidin-1-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
356	3-amino-N-methyl-6-[3-(((2-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
357	1,1-dimethylethyl (3R)-3-(((3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]phenyl)carbonyl)amino)pyrrolidine-1-carboxylate
358	3-amino-N-methyl-6-(3-(((3R)-pyrrolidin-3-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide
359	3-(((4-chlorophenyl)methyl)amino)-6-[3-(((3R)-1-[(4-chlorophenyl)methyl]pyrrolidin-3-yl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
360	3-amino-6-[3-(((3R)-1-[(2,4-dichlorophenyl)methyl]pyrrolidin-3-yl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
361	N-methyl-3-(((3-(methyloxy)phenyl)methyl)amino)-6-[3-(((3R)-1-[(3-(methyloxy)phenyl)methyl]pyrrolidin-3-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
362	3-amino-6-[3-(((3R)-1-[(2,6-dichlorophenyl)methyl]pyrrolidin-3-yl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
363	N-methyl-3-[(phenylmethyl)amino]-6-[3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
364	1,1-dimethylethyl 3-(((3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]phenyl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
365	1,1-dimethylethyl 2-(((3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]phenyl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
366	3-amino-N-methyl-6-(3-(((pyrrolidin-3-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
367	3-amino-N-methyl-6-(3-(((pyrrolidin-2-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
368	3-amino-6-(2-fluoro-5-(((phenylmethyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
369	3-amino-6-(3-(((1,3-benzodioxol-5-ylacetyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
370	3-amino-6-[3-(((2-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
371	3-amino-6-[3-(((3,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
372	3-amino-N-methyl-6-[3-[(phenylmethyl)amino]phenyl]pyrazine-2-carboxamide
373	3-amino-N-methyl-6-(3-(((2-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide

#	Name
374	3-amino-N-methyl-6-{3-[[[4-(1-methylethyl)phenyl]carbonyl]amino)methyl]phenyl}pyrazine-2-carboxamide
375	3-amino-6-[3-[[[4-butylphenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
376	3-amino-N-methyl-6-{3-[[[4-(1,2,3-thiadiazol-4-yl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
377	3-amino-N-methyl-6-{3-[[[4-(2-thienyl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
378	3-amino-6-[3-[[[4-(1,1-dimethylethyl)phenyl]carbonyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
379	3-amino-N-methyl-6-{3-[[[4-(1H-pyrazol-1-yl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
380	3-amino-6-[3-[[[3,4-dihydro-2H-1,4-benzoxazin-6-yl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
381	3-amino-6-[3-[[[2-amino-6-fluorophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
382	3-amino-6-[3-[[[biphenyl-4-yl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
383	3-amino-6-[3-[[[biphenyl-2-yl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
384	3-amino-6-[3-[[[biphenyl-3-yl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
385	3-amino-N-methyl-6-{3-[[[1-(phenylmethyl)pyrrolidin-2-yl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
386	3-amino-6-[3-[[[3,4-dihydro-2H-1,4-benzoxazin-6-yl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
387	3-amino-6-[3-[[[2,4-difluorophenyl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
388	3-amino-6-[3-[[[2,5-difluorophenyl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
389	3-amino-6-[3-[[[2,3-difluorophenyl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
390	3-amino-N-methyl-6-[3-[[[2-pyridin-4-ylphenyl)methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
391	3-amino-N-methyl-6-[3-[[[2-pyridin-3-ylphenyl)methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
392	3-amino-N-methyl-6-[3-[[[4-(methylsulfonyl)phenyl]methyl]amino]methyl]phenyl}pyrazine-2-carboxamide
393	3-amino-6-[3-[[[2,2-diphenylethyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide
394	3-amino-N-methyl-6-[3-[[[1-(phenylmethyl)pyrrolidin-3-yl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
395	3-amino-N-methyl-6-[3-[[[3-(phenyloxy)phenyl]methyl]amino]methyl]phenyl}pyrazine-2-carboxamide
396	3-amino-6-[3-[[[4-aminophenyl)methyl]amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
397	3-amino-N-methyl-6-[3-(((4-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
398	3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
399	3-amino-6-[3-(((2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
400	3-amino-6-[5-(((2,6-difluorophenyl)methyl)amino)carbonyl]pyridin-3-yl]-N-methylpyrazine-2-carboxamide
401	3-amino-6-(5-(((biphenyl-2-ylmethyl)amino)carbonyl]pyridin-3-yl)-N-methylpyrazine-2-carboxamide
402	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
403	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
404	3-amino-N-methyl-6-[3-(((2-piperidin-1-ylphenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
405	3-amino-6-[3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
406	3-amino-N-methyl-6-[3-(((6-(trifluoromethyl)pyridin-3-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
407	3-amino-6-[3-(1H-benzimidazol-2-yl)phenyl]-N-methylpyrazine-2-carboxamide
408	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
409	3-amino-6-[3-(((3,5-dichlorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
410	3-amino-N-methyl-6-[3-((2-phenylethyl)amino)phenyl]pyrazine-2-carboxamide
411	3-amino-6-(5-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl]pyridin-3-yl)-N-methylpyrazine-2-carboxamide
412	3-amino-N-methyl-6-[3-(((2-(phenylamino)ethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
413	3-amino-6-[3-(((4-ethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
414	3-amino-N-methyl-6-[3-(((4-pyridin-3-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
415	3-amino-6-[3-((2,3-dihydro-1H-inden-2-ylamino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
416	3-amino-6-[3-(1,3-dihydro-2H-isoindol-2-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide
417	3-amino-6-[3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
418	3-amino-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
419	3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide

#	Name
420	3-amino-N-methyl-6-[3-(((1R)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide
421	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
422	3-amino-N-methyl-6-[3-(((4-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
423	3-amino-N-methyl-6-[3-(((3-(1,2,3-thiadiazol-4-yl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
424	3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
425	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
426	3-amino-N-methyl-6-[3-(((3-(trifluoromethyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
427	3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide
428	3-amino-6-[3-(((1R)-2-hydroxy-1-phenylethyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
429	methyl (2S)-(((3-(5-amino-6-((methylamino)carbonyl]pyrazin-2-yl)phenyl)methyl)amino)(phenyl)ethanoate
430	3-amino-6-[3-([2-(2,5-difluorophenyl)hydrazino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
431	3-amino-N-methyl-6-[3-([2-(2,3,5,6-tetrafluoro-4-methylphenyl)hydrazino]carbonyl)phenyl]pyrazine-2-carboxamide
432	3-amino-6-[3-([2-(2-fluorophenyl)hydrazino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
433	3-amino-6-[3-([2-(2,4-difluorophenyl)hydrazino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
434	3-amino-N-methyl-6-[3-([2-(2,3,5,6-tetrafluorophenyl)hydrazino]carbonyl)phenyl]pyrazine-2-carboxamide
435	3-amino-N-methyl-6-[3-(((5-methylpyrazin-2-yl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
436	3-amino-N-methyl-6-[3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)methyl)phenyl]pyrazine-2-carboxamide
437	3-amino-N-methyl-6-[3-(((3S)-1-(phenylmethyl)pyrrolidin-3-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
438	3-amino-N-methyl-6-[3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
439	3-amino-6-[3-([ethyl(phenylmethyl)amino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
440	3-amino-N-methyl-6-[3-(((1S,2S)-2-phenylcyclopropyl)amino)methyl]phenyl]pyrazine-2-carboxamide
441	3-Amino-6-(3-benzylcarbonyl-phenyl)-pyrazine-2-carboxylic acid methylamide
442	3-amino-N-methyl-6-[3-(4-phenyl-1H-imidazol-2-yl)phenyl]pyrazine-2-carboxamide

#	Name
443	3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
444	3-amino-6-[3-(((5-bromo-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
445	3-amino-N-methyl-6-(3-(((2R)-1,2,3,4-tetrahydronaphthalen-2-ylamino)methyl)phenyl)pyrazine-2-carboxamide
446	3-amino-N-methyl-6-(3-(((1-methyl-1-phenylethyl)amino)methyl)phenyl)pyrazine-2-carboxamide
447	3-amino-6-[3-((9H-fluoren-9-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
448	3-amino-N-methyl-6-[3-((naphthalen-1-ylamino)methyl)phenyl]pyrazine-2-carboxamide
449	3-amino-6-[3-(((3-fluorobiphenyl-4-yl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
450	3-amino-6-[3-(((2-fluoro-4-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
451	3-amino-6-[3-(((2-fluoro-4-furan-2-ylphenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
452	3-amino-N-methyl-6-[3-((naphthalen-2-ylamino)methyl)phenyl]pyrazine-2-carboxamide
453	3-amino-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
454	3-amino-N-methyl-6-(3-(((2S)-1,2,3,4-tetrahydronaphthalen-2-ylamino)methyl)phenyl)pyrazine-2-carboxamide
455	methyl 3-amino-6-(3-(((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)pyrazine-2-carboxylate
456	3-amino-6-(3-(((5-bromo-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
457	3-amino-6-[3-(((2-fluoro-5-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
458	3-amino-6-[3-(((4-ethenyl-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
459	3-amino-6-(3-(((4,7-difluoro-3-methyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
460	3-amino-N-methyl-6-[3-(((1S,2R)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
461	3-amino-6-[3-(((5-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
462	3-amino-6-[3-(((2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
463	3-amino-6-[3-(((2R)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
464	3-amino-6-(3-(((5-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
465	3-amino-6-[3-(((2S)-5-bromo-2,3-dihydro-1H-inden-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
466	3-amino-6-(3-(((4-fluoro-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
467	3-amino-6-(3-(((4-fluoro-1-hydroxy-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
468	3-amino-N-methyl-6-[3-(((1R,2S)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
469	3-amino-6-(3-(((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
470	3-amino-6-(3-(((6-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
471	3-amino-N-methyl-6-(3-(((3-oxo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)pyrazine-2-carboxamide
472	3-amino-6-(3-(((5-furan-2-yl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
473	3-amino-6-(3-(((4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
474	3-amino-6-(3-(((4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
475	3-amino-6-(3-(((6-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
476	3-amino-6-(3-(((5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
477	3-amino-6-(3-(((5-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
478	3-amino-N-methyl-6-[3-(((5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
479	3-amino-6-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
480	3-amino-6-(3-(((5-furan-3-yl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
481	3-amino-N-methyl-6-[3-(((5-(3-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
482	3-amino-6-[3-(((1S)-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
483	3-amino-6-[3-(((1S)-5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
484	3-amino-6-(3-(((7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
485	3-amino-6-(3-(((6-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
486	3-amino-6-(3-(((4,6-dichloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
487	3-amino-6-(3-(((4-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
488	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-methylpyrazine-2-carboxamide

#	Name
489	3-amino-6-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-ethylpyrazine-2-carboxamide
490	3-amino-N-methyl-6-[3-(((1S)-5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
491	3-amino-N-methyl-6-(3-(((5-phenyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)pyrazine-2-carboxamide
492	3-amino-6-(3-(((4-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
493	3-amino-N-methyl-6-(3-(((4-phenyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)pyrazine-2-carboxamide
494	3-amino-N-methyl-6-[3-(((4-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
495	3-amino-N-methyl-6-[3-(((6-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
496	3-amino-N-methyl-6-[3-(((5-(4-methyl-2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
497	3-amino-N-methyl-6-[3-(((2R)-6-phenyl-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
498	3-amino-N-methyl-6-naphthalen-2-ylpyrazine-2-carboxamide
499	3-amino-N-methyl-6-[3-(((1S,2S)-2-((phenylmethyl)oxy)cyclopentyl)amino)methyl]phenyl]pyrazine-2-carboxamide
500	3-amino-6-(3-((cyclopentylamino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
501	3-amino-6-(3-(((6-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
502	3-amino-6-(3-(((7-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
503	3-amino-N-methyl-6-quinolin-3-ylpyrazine-2-carboxamide
504	3-amino-6-[3-(1H-imidazol-1-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide
505	3-amino-N-methyl-6-[3-(((5-[(2-morpholin-4-ylethyl)oxy]-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
506	3-amino-N-methyl-6-[3-(((1R,2R)-2-((phenylmethyl)oxy)cyclopentyl)amino)methyl]phenyl]pyrazine-2-carboxamide
507	3-amino-6-[3-((2,3-dihydro-1-benzofuran-3-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
508	3-amino-6-[3-(((5-[(cyanomethyl)oxy]-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
509	2-amino-5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-methylpyridine-3-carboxamide
510	3-amino-N-methyl-6-[3-(((5-(4-methylphenyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
511	3-amino-6-[3-(((2-(dimethylamino)-1-phenylethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide

#	Name
512	3-amino-N-methyl-6-[3-((5-(1H-1,2,4-triazol-1-yl)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
513	3-amino-6-(3-((5-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
514	3-amino-N-methyl-6-[3-((4-((2-morpholin-4-ylethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
515	3-amino-6-(3-((4-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
516	3-amino-N-methyl-6-{1-[(phenylmethyl)amino]isoquinolin-7-yl}pyrazine-2-carboxamide
517	3-amino-N-methyl-6-(3-((2-morpholin-4-yl-1-phenylethyl)amino)methyl)phenyl]pyrazine-2-carboxamide
518	3-amino-N-methyl-6-(3-((1-phenyl-2-pyrrolidin-1-ylethyl)amino)methyl)phenyl]pyrazine-2-carboxamide
519	2-amino-5-(3-((6-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
520	2-amino-5-(3-((4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
521	2-amino-5-(3-((7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
522	2-amino-5-(3-((4-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
523	2-amino-5-(3-((5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
524	2-amino-5-(3-((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
525	3-amino-6-{1-[(1S)-2,3-dihydro-1H-inden-1-ylamino]isoquinolin-7-yl}-N-methylpyrazine-2-carboxamide
526	3-amino-6-[3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]-2,3-dihydro-1H-inden-5-yl]-N-methylpyrazine-2-carboxamide
527	3-amino-6-[3-((6-(3-hydroxypropyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
528	5-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(1,3-oxazol-5-yl)pyrazin-2-amine
529	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]pyrazine-2-carbohydrazide
530	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]pyrazine-2-carboxamide
531	1-[3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]pyrazin-2-yl]ethanone
532	3-amino-6-[3-[(2,3-dihydro-1H-inden-1-ylamino)methyl]phenyl]-N-(naphthalen-2-ylmethyl)pyrazine-2-carboxamide
533	3-amino-N-cyclohexyl-6-[3-[(2,3-dihydro-1H-inden-1-ylamino)methyl]phenyl]pyrazine-2-carboxamide
534	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(pyridin-4-ylmethyl)pyrazine-2-carboxamide

#	Name
535	{3-[5-amino-6-(4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol
536	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(4H-1,2,4-triazol-3-yl)pyrazin-2-amine
537	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2,2,2-trifluoroethyl)pyrazine-2-carboxamide
538	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(5-hydroxypentyl)pyrazine-2-carboxamide
539	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(4-hydroxybutyl)pyrazine-2-carboxamide
540	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-pyrrolidin-1-ylethyl)pyrazine-2-carboxamide
541	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-piperidin-1-ylethyl)pyrazine-2-carboxamide
542	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-morpholin-4-ylethyl)pyrazine-2-carboxamide
543	3-amino-N-(cyclopropylmethyl)-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
544	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(phenylmethyl)pyrazine-2-carboxamide
545	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-phenylethyl)pyrazine-2-carboxamide
546	[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazin-2-yl](phenyl)methanone
547	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-hydroxyethyl)pyrazine-2-carboxamide
548	3-amino-N-cyclopropyl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
549	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine
550	{3-[5-amino-6-(5-methyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol
551	{3-[5-amino-6-(5-phenyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol
552	(3-{5-amino-6-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol
553	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N'-methylpyrazine-2-carbohydrazide
554	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N,N-dimethylpyrazine-2-carboxamide
555	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
556	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(dimethylamino)ethyl]pyrazine-2-carboxamide
557	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-phenyl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine

#	Name
558	3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide
559	3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[2-(methylamino)ethyl]pyrazine-2-carboxamide
560	3-{5-amino-6-[(3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
561	3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-(3-hydroxypropyl)pyrazine-2-carboxamide
562	1,1-dimethylethyl 4-([(3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)piperidine-1-carboxylate
563	1,1-dimethylethyl 4-([(3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)methyl]piperidine-1-carboxylate
564	3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)-N-piperidin-4-ylpyrazine-2-carboxamide
565	3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)-N-(piperidin-4-ylmethyl)pyrazine-2-carboxamide
566	3-[5-amino-6-(morpholin-4-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
567	1,1-dimethylethyl 3-([(3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)pyrrolidine-1-carboxylate
568	3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)-N-pyrrolidin-3-ylpyrazine-2-carboxamide
569	(3-{5-amino-6-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol
570	{3-[5-amino-6-(5-furan-2-yl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl)methanol
571	[3-(5-amino-6-[5-[3-(methyloxy)phenyl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)phenyl)methanol
572	2-([(3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl)carbonyl)-N-phenylhydrazinecarboxamide
573	3-{5-amino-6-[(4-methylpiperazin-1-yl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
574	3-(5-amino-6-[[4-(4-fluorophenyl)piperazin-1-yl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
575	3-(5-amino-6-[[4-(phenylmethyl)piperazin-1-yl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
576	methyl N-[(3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl]glycinate
577	N-[(3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl]glycine
578	3-{5-amino-6-[(3,5-difluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
579	3-[5-amino-6-(biphenyl-4-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide

#	Name
580	{3-[5-amino-6-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl)methanol
581	(3-{5-amino-6-[3-(4-chlorophenyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol
582	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
583	(3-{5-amino-6-[3-(2-thienyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol
584	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-[3-(methyloxy)phenyl]-1H-1,2,4-triazol-5-yl]pyrazin-2-amine
585	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[3-(dimethylamino)propyl]pyrazine-2-carboxamide
586	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[3-(1H-imidazol-1-yl)propyl]pyrazine-2-carboxamide
587	3-{5-amino-6-[(4-chloro-3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
588	3-(5-amino-6-[[2,4-bis(methyloxy)phenyl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
589	3-(5-amino-6-[[4-(dimethylamino)phenyl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
590	methyl 1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]-L-prollinate
591	methyl 1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]piperidine-3-carboxylate
592	1,1-dimethylethyl 4-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]piperazine-1-carboxylate
593	3-[5-amino-6-(piperazin-1-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
594	N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]glutamic acid
595	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-[5-[[[(methyloxy)methyl]-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
596	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
597	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[3-(methylamino)propyl]pyrazine-2-carboxamide
598	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-(3-pyrrolidin-1-ylpropyl)pyrazine-2-carboxamide
599	3-(5-amino-6-[(1E)-N-[4-(methyloxy)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
600	3-(5-amino-6-[(1E)-N-[4-(1-methylethyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
601	3-{5-amino-6-[(1E)-N-1,3-benzothiazol-2-ylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
602	3-{5-amino-6-[(1E)-N-(4-methylphenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide

#	Name
603	3-{5-amino-6-[(1E)-N-(4-chlorophenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
604	3-{5-amino-6-[(1E)-N-methyl-N-phenylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
605	3-{5-amino-6-[(1E)-N-(2-hydroxyethyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
606	{3-[5-amino-6-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
607	{3-[5-amino-6-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
608	3-[5-amino-6-(2-aminopyrimidin-4-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
609	3-{5-amino-6-[2-(dimethylamino)pyrimidin-4-yl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
610	ethyl N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]-L-tyrosinate
611	N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]-D-tyrosine
612	1,1-dimethylethyl [3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]propyl]carbamate
613	3-amino-N-(3-aminopropyl)-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
614	1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]piperidine-3-carboxylic
615	1,1-dimethylethyl [2-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]ethyl]carbamate
616	3-amino-N-(2-aminoethyl)-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
617	1,1-dimethylethyl (1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]piperidin-4-yl]carbamate
618	3-{5-amino-6-[(4-aminopiperidin-1-yl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
619	3-{5-amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
620	3-{5-amino-6-[imino(morpholin-4-ylamino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
621	3-(5-amino-6-[imino[(4-methylpiperazin-1-yl)amino]methyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
622	3-{5-amino-6-[imino(piperidin-1-ylamino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
623	3-{5-amino-6-[(azepan-1-ylamino)(imino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
624	3-{5-amino-6-[imino((2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl)amino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide

#	Name
625	{3-[5-amino-6-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
626	3-[5-amino-6-[(1E)-N-morpholin-4-ylethanimidoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
627	4-((2E)-2-{1-[3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl]pyrazin-2-yl}ethylidene)hydrazino)benzoic acid
628	ethyl ((2E)-2-{1-[3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl]pyrazin-2-yl}ethylidene)hydrazino)acetate
629	3-[5-amino-6-[(1E)-N,N-dimethylethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
630	3-(5-amino-6-[(1E)-N-[4-(methylsulfonyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
631	3-[5-amino-6-[(1E)-N-(4-cyanophenyl)ethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
632	3-[5-amino-6-[(1E)-N-pyridin-2-ylethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
633	3-(5-amino-6-[(1E)-N-[amino(imino)methyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
634	3-[5-amino-6-[(1E)-N-[4-[(trifluoromethyl)oxy]phenyl]ethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
635	3-(5-Amino-6-{1-[(4-nitro-phenyl)-hydrazono]-ethyl}-pyrazin-2-yl)-N-benzylbenzamide
636	3-(5-amino-6-[(1E)-N-[4-(trifluoromethyl)pyrimidin-2-yl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
637	3-[5-amino-6-[(1E)-N-1H-1,2,3-benzotriazol-1-ylethanimidoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
638	3-[5-amino-6-[(1E)-N-methylethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
639	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
640	3-(5-amino-6-[(1E)-N-[4-(trifluoromethyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
641	3-[5-amino-6-[(1E)-N-phenylethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
642	3-[5-amino-6-[(1E)-N-(4-methylpiperazin-1-yl)ethanimidoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
643	{3-[5-amino-6-(3-pyrrolidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
644	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
645	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-yl]pyrazine-2-carboxamide
646	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(furan-2-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide
647	N-[3-(acetylamino)propyl]-3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide

#	Name
648	3-amino-N-[3-[(furan-2-ylcarbonyl)amino]propyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
649	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(pyridin-4-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide
650	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(pyridin-4-ylcarbonyl)amino]propyl]pyrazine-2-carboxamide
651	N-[3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]propyl]quinoxaline-2-carboxamide
652	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[[[6-(trifluoromethyl)pyridin-3-yl]carbonyl]amino]propyl]pyrazine-2-carboxamide
653	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(quinolin-8-ylsulfonyl)amino]propyl]pyrazine-2-carboxamide
654	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(pyridin-2-ylsulfonyl)amino]propyl]pyrazine-2-carboxamide
655	3-amino-N-[3-[[[(3,5-dimethylisoxazol-4-yl)amino]carbonyl]amino]propyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
656	3-[5-amino-6-(2-methylpyrimidin-4-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
657	3-[5-amino-6-[2-(methylamino)pyrimidin-4-yl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
658	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[2-[(pyridin-4-ylcarbonyl)amino]ethyl]pyrazine-2-carboxamide
659	3-amino-N-[2-[(furan-2-ylcarbonyl)amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
660	N,N'-cyclohexane-1,2-diylbis[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide]
661	N-[2-(acetylamino)ethyl]-3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
662	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[2-[(quinolin-8-ylsulfonyl)amino]ethyl]pyrazine-2-carboxamide
663	N-[2-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]ethyl]quinoxaline-2-carboxamide
664	3-amino-N-[2-[[[2-chloropyridin-3-yl]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
665	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[2-[(pyridin-2-ylsulfonyl)amino]ethyl]pyrazine-2-carboxamide
666	3-amino-N-[2-[[[(3,5-dimethylisoxazol-4-yl)amino]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
667	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
668	3-[5-amino-6-(imino[2-[4-(trifluoromethyl)pyrimidin-2-yl]hydrazino)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
669	3-[5-amino-6-[2-(1,3-benzothiazol-2-yl)hydrazino](imino)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
670	3-[5-amino-6-(1,5-diphenyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide

#	Name
671	3-[5-amino-6-(1,5-dimethyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
672	3-[5-amino-6-(1-methyl-5-phenyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
673	3-[5-amino-6-((1E)-N-((2R)-2-((methyloxy)methyl)pyrrolidin-1-yl)ethanimidoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
674	3-[5-amino-6-((1E)-N-azepan-1-ylethanimidoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
675	3-(5-amino-6-((E)-[(phenylmethyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
676	3-[5-amino-6-((E)-[amino(imino)methyl]hydrazono)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
677	3-(5-amino-6-((E)-[(2-hydroxyethyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
678	3-(5-amino-6-((E)-(pyridin-2-ylhydrazono)methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
679	3-(5-amino-6-((E)-[(4-cyanophenyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
680	3-(5-amino-6-((E)-[(4-methylphenyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
681	3-(5-amino-6-((E)-(hydroxyimino)methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
682	{3-[5-amino-6-(3-pyridin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
683	5-(3-[[[(1R)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
684	3-amino-N-[3-[[[4-(dimethylamino)phenyl]carbonyl]amino]propyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
685	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
686	1,1-dimethylethyl 4-[(3S)-3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidin-1-yl]piperidine-1-carboxylate
687	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[(3S)-1-piperidin-4-ylpyrrolidin-3-yl]pyrazine-2-carboxamide
688	1,1-dimethylethyl (2R)-2-[[[(3S)-3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidin-1-yl]methyl]pyrrolidine-1-carboxylate
689	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[(3S)-1-[(2R)-pyrrolidin-2-ylmethyl]pyrrolidin-3-yl]pyrazine-2-carboxamide
690	3-amino-N-((3S)-1-[[[4-(dimethylamino)phenyl]methyl]pyrrolidin-3-yl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
691	3-amino-N-[2-[[[4-(dimethylamino)phenyl]amino]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
692	3-amino-N-[2-[[[4-(dimethylamino)phenyl]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
693	3-(5-amino-6-[imino(2-pyridin-2-ylhydrazino)methyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide

#	Name
694	3-[5-amino-6-(morpholin-4-ylmethyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
695	ethyl 1-[[3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl]pyrazin-2-yl]methyl]piperidine-4-carboxylate
696	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[(3S)-1-(pyridin-2-ylmethyl)pyrrolidin-3-yl]pyrazine-2-carboxamide
697	methyl N-[[3-amino-6-(3-[(phenylmethyl)amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl]threoninate
698	{3-[5-amino-6-(3-piperidin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
699	3-(ethylamino)-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-(3-[(phenylmethyl)amino]carbonyl)phenyl]pyrazine-2-carboxamide
700	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-(3-[(phenylmethyl)amino]carbonyl)phenyl]pyrazine-2-carboxamide
701	3-[5-amino-6-(1-methyl-5-piperidin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
702	3-[5-amino-6-(5-methyl-1-pyridin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
703	{3-[5-amino-6-(3-morpholin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
704	5-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-3-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine
705	5-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-3-{3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine
706	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-N-[(3S)-1-ethylpyrrolidin-3-yl]pyrazine-2-carboxamide
707	6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-3-(ethylamino)-N-[(3S)-1-ethylpyrrolidin-3-yl]pyrazine-2-carboxamide
708	3-amino-6-[3-[(1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
709	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
710	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[(1R)-1-phenylethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
711	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-(3,4-dihydroisoquinolin-2(1H)-ylcarbonyl)phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
712	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(2-phenylethyl)amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
713	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[(1S)-1-phenylethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
714	3-amino-6-[3-(3,4-dihydroisoquinolin-2(1H)-ylcarbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
715	3-amino-6-(3-[(2-phenylethyl)amino]carbonyl)phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide

#	Name
716	3-amino-6-[3-(((1R)-1-phenylethyl)amino)carbonyl]phenyl-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
717	3-amino-6-[3-(((1S)-1-phenylethyl)amino)carbonyl]phenyl-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
718	3-{5-amino-6-[(2-ethylhydrazino)(imino)methyl]pyrazin-2-yl}-N-(2,3-dihydro-1H-inden-1-yl)benzamide
719	3-{5-amino-6-[imino(2-methylhydrazino)methyl]pyrazin-2-yl}-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
720	3-{5-amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl}-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
721	phenylmethyl 3-{3-[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl}piperidine-1-carboxylate
722	phenylmethyl 3-{3-[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl}piperidine-1-carboxylate
723	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
724	[3-(5-amino-6-{3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methanol
725	[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methanol
726	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
727	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
728	1,1-dimethylethyl 3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
729	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)-N-piperidin-3-ylpyrazine-2-carboxamide
730	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)-N-(1-ethylpiperidin-3-yl)pyrazine-2-carboxamide
731	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
732	(3-{5-amino-6-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol
733	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
734	3-[5-amino-6-(1-ethyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]benzamide
735	3-[5-amino-6-(1-methyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
736	(3-{5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol
737	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-amine

#	Name
738	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-(methoxy)phenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
739	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
740	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[2-chloro-6-fluorophenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
741	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-(methylthio)phenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
742	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
743	3-amino-6-{3-[[[4-(methoxy)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
744	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-{3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide
745	3-amino-6-{3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
746	3-amino-6-{3-[[[2-chloro-6-fluorophenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
747	3-amino-6-{3-[[[4-(methylthio)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
748	3-{5-amino-6-[imino(2-pyridin-4-ylhydrazino)methyl]pyrazin-2-yl}-N-[(1R)-2,3-dihydro-1H-inden-1-yl]benzamide
749	phenylmethyl (3R)-3-{3-[3-amino-6-{3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl}pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl}piperidine-1-carboxylate
750	3-amino-6-{3-[(1S)-1-[(1S)-2,3-dihydro-1H-inden-1-ylamino]ethyl]phenyl}-N-methylpyrazine-2-carboxamide
751	3-amino-6-{3-[[[3-[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide
752	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]methyl]pyrrolidine-1-carboxylate
753	3-amino-6-{3-[[[3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
754	phenylmethyl (3R)-3-{3-[3-amino-6-{3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl}pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl}piperidine-1-carboxylate
755	3-{5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl}-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
756	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-chlorophenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
757	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[(pyridin-2-ylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
758	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[(pyridin-3-ylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate

#	Name
759	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-((3,4-dihydro-2H-chromen-4-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
760	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
761	3-amino-6-(3-((pyridin-2-ylmethyl)amino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
762	3-amino-6-(3-((pyridin-3-ylmethyl)amino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
763	dimethyl 5-(5-amino-6-((methylamino)carbonyl)pyrazin-2-yl)benzene-1,3-dicarboxylate
764	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
765	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-[(3R)-piperidin-3-yl]pyrazine-2-carboxamide
766	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-piperidin-3-ylpyrazine-2-carboxamide
767	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide
768	3-amino-N-azepan-4-yl-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]pyrazine-2-carboxamide
769	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
770	1,1-dimethylethyl 4-(((3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate
771	3-[5-amino-6-(5-(2-((phenylmethyl)oxy)ethyl)-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
772	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)-4-fluorophenyl)-N-methylpyrazine-2-carboxamide
773	3-[5-amino-6-(1-phenyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
774	3-[5-amino-6-[amino(imino)methyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
775	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]benzamide
776	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
777	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
778	3-amino-6-[3-((3,4-dihydro-2H-chromen-4-ylamino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
779	3-amino-6-[3-((2,3-dihydro-1-benzofuran-3-ylamino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
780	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
781	1,1-dimethylethyl 4-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate

#	Name
782	3-amino-N-azepan-4-yl-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
783	3-amino-N-azepan-4-yl-6-[3-[(3aR,8aS)-8,8a-dihydro-3aH-indeno[1,2-d][1,3]oxazol-2-yl]phenyl]pyrazine-2-carboxamide
784	1,1-dimethylethyl (3R)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
785	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
786	1,1-dimethylethyl (3R)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
787	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
788	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)-5-fluorophenyl)-N-methylpyrazine-2-carboxamide
789	3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-5-[(methyloxy)carbonyl]benzoic acid
790	methyl 3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-5-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]benzoate
791	5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-N'-pyrrolidin-3-ylbenzene-1,3-dicarboxamide
792	3-[5-amino-6-(5-phenyl-1-pyridin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
793	3-[5-amino-6-(5-methyl-1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
794	1,1-dimethylethyl (3R)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
795	1,1-dimethylethyl (3R)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
796	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
797	5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-N'-[(3R)-pyrrolidin-3-yl]benzene-1,3-dicarboxamide
798	(3-[5-amino-6-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl]phenyl)methanol
799	3-[5-amino-6-(1,5-dimethyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
800	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-[(1-methyl-1-phenylethyl)amino]carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
801	3-amino-6-(3-[(1-methyl-1-phenylethyl)amino]carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
802	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)sulfonyl]phenyl)pyrazine-2-carboxamide
803	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)sulfonyl]phenyl)-N-piperidin-3-ylpyrazine-2-carboxamide

#	Name
804	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-((3S)-1-methylpyrrolidin-3-yl)pyrazine-2-carboxamide
805	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-((3S)-1-methylpyrrolidin-3-yl)pyrazine-2-carboxamide
806	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
807	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-((3S)-piperidin-3-yl)pyrazine-2-carboxamide
808	3-amino-N-((3S)-pyrrolidin-3-yl)-6-(3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
809	N-[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl})pyrazin-2-yl]phenyl]-2-(2,4-dichlorophenyl)acetamide
810	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)sulfonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
811	3-amino-6-(3-hydroxyphenyl)-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
812	3-amino-6-(3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)sulfonyl)phenyl)-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
813	3-[5-amino-6-(1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-((1S)-2,3-dihydro-1H-inden-1-yl)benzamide
814	3-amino-N-((3S)-pyrrolidin-3-yl)-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
815	3-amino-N-((3S)-pyrrolidin-3-yl)-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
816	3-amino-N-((3S)-pyrrolidin-3-yl)-6-[3-(((2,3,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
817	3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
818	3-(5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2-chloro-6-fluorophenyl)methyl]benzamide
819	3-amino-N-1-azabicyclo[2.2.2]oct-3-yl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
820	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(4-chloro-2-fluorophenyl)methyl]benzamide
821	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2-chlorophenyl)methyl]benzamide
822	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
823	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
824	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(4-piperidin-3-yl-1,3-thiazol-2-yl)pyrazin-2-amine
825	3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide

#	Name
826	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
827	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2,6-difluorophenyl)methyl]benzamide
828	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2-chloro-6-fluorophenyl)methyl]benzamide
829	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2,5-dichlorophenyl)methyl]benzamide
830	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(3,4-dichlorophenyl)methyl]benzamide
831	3-amino-6-[3-[[[4-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
832	3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
833	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[[[1,1-dimethylethyl]oxy]carbonyl]amino)methyl]phenyl]pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate
834	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(2-fluorophenyl)methyl]benzamide
835	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(3-fluorophenyl)methyl]benzamide
836	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(4-fluorophenyl)methyl]benzamide
837	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(3-chlorophenyl)methyl]benzamide
838	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(4-chlorophenyl)methyl]benzamide
839	3-amino-6-[3-(aminomethyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
840	3-amino-6-(3-hydroxyphenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
841	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate
842	3-amino-6-[3-(((4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
843	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
844	3-amino-6-[3-[[[biphenyl-4-ylmethyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
845	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-(1,2,3-thiadiazol-5-yl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
846	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[2-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
847	3-amino-6-[3-[[[biphenyl-2-ylmethyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
848	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
849	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
850	3-amino-6-[3-(((4-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
851	3-amino-6-[3-(((naphthalen-2-ylmethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
852	3-amino-6-[3-(((4-(dimethylamino)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
853	3-amino-6-[3-(((1,3-benzodioxol-5-ylmethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
854	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(2-thienyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
855	3-amino-6-[3-(((2-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
856	3-amino-6-[3-(((2,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
857	3-amino-6-[3-(((4-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
858	3-amino-6-[3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
859	3-amino-6-[3-(((4-(methylsulfonyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
860	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
861	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
862	3-amino-6-[3-(((5-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
863	3-amino-N-[(3R)-1-azabicyclo[2.2.2]oct-3-yl]-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]pyrazine-2-carboxamide
864	3-amino-6-[3-(((4-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
865	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2,4-difluorophenyl)methyl]benzamide
866	3-amino-6-[3-(((1,2-diphenylethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
867	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(3,4-difluorophenyl)methyl]benzamide
868	3-amino-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
869	methyl 4-(((3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl]pyrazin-2-yl)phenyl)carbonyl)amino)methylbenzoate
870	3-amino-6-[3-(((4-bromophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
871	3-amino-6-[3-(((4-bromo-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
872	3-amino-6-[3-(((2-bromophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
873	3-amino-6-[3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
874	3-amino-6-[3-(((2-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
875	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3R)-pyrrolidin-3-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
876	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((3R)-pyrrolidin-3-ylamino)carbonyl]phenyl)pyrazine-2-carboxamide
877	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-((2-piperidin-1-ylethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
878	3-amino-6-(3-((2-phenylethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
879	3-amino-6-[3-((2-(methyloxy)ethyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
880	3-amino-6-[3-(((4-iodophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
881	3-amino-6-[3-(((2-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
882	3-amino-6-[3-(((4-fluorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
883	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((4-((trifluoromethyl)oxy)phenyl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide
884	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
885	3-amino-6-(3-((4-chlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
886	3-amino-6-(3-((2,6-dichlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
887	3-amino-6-(3-((pentafluorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
888	3-amino-6-(3-((2-chloro-4-fluorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
889	3-amino-6-[3-(((4-(methylthio)phenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
890	3-amino-6-(3-((2,4-dichlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
891	3-amino-6-[3-(((4-(4-chlorophenyl)-2-thienyl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
892	3-amino-6-[3-(((4-bromophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
893	3-amino-6-(3-(((naphthalen-2-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
894	3-amino-6-(3-([2-(4-phenylpiperazin-1-yl)ethyl]oxy)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
895	N-[(3-(5-amino-6-[(3R)-piperidin-3-ylacetyl]pyrazin-2-yl)phenyl)methyl]-N'-(4-bromophenyl)urea
896	N-[(3-(5-amino-6-[(3R)-piperidin-3-ylacetyl]pyrazin-2-yl)phenyl)methyl]-N'-naphthalen-2-ylurea
897	3-amino-6-(3-(((2,4-difluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
898	3-amino-6-(3-(((2-fluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
899	3-amino-6-(3-(((5-[2-chloro-5-(trifluoromethyl)phenyl]furan-2-yl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
900	3-amino-6-[3-(((4-(dimethylamino)naphthalen-1-yl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
901	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1,2,3-thiadiazol-5-yl)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide
902	3-amino-6-(3-(((1,3-dimethyl-1H-pyrazol-5-yl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
903	3-amino-6-(3-(((4-(dimethylamino)phenyl)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
904	3-amino-6-(3-([2-(hydroxyethyl)oxy]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
905	3-amino-6-(3-([2,3-dihydro-1-benzofuran-5-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
906	3-amino-6-(3-([3,4-dihydro-2H-1,5-benzodioxepin-7-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
907	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-([quinolin-7-ylsulfonyl)amino)methyl]phenyl]pyrazine-2-carboxamide
908	3-amino-6-[3-([[(biphenyl-4-ylamino)carbonyl]amino)methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
909	3-amino-6-(3-([4-bromo-2-fluorophenyl)carbonyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
910	N-[3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl)pyrazin-2-yl]phenyl]quinoline-3-carboxamide
911	3-amino-6-[3-([6-(methyloxy)-1-benzofuran-3-yl]acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
912	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-([4'-(trifluoromethyl)biphenyl-2-yl]carbonyl)amino)phenyl]pyrazine-2-carboxamide
913	3-amino-6-[3-([3,5-bis(trifluoromethyl)phenyl]acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
914	3-amino-6-[3-([[(2,4-difluorophenyl)amino]carbonyl)amino)methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
915	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-([2-(trifluoromethyl)phenyl]sulfonyl)amino)methyl]phenyl]pyrazine-2-carboxamide

#	Name
916	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[4-[(trifluoromethyl)oxy]phenyl)sulfonyl]amino]methyl]phenyl)pyrazine-2-carboxamide
917	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[2-[(trifluoromethyl)oxy]phenyl)sulfonyl]amino]methyl]phenyl)pyrazine-2-carboxamide
918	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[4-(trifluoromethyl)phenyl)sulfonyl]amino]methyl]phenyl)pyrazine-2-carboxamide
919	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[2-[(trifluoromethyl)oxy]phenyl]amino]carbonyl]amino]methyl]phenyl)pyrazine-2-carboxamide
920	3-amino-6-(3-[[[2-(methylthio)phenyl]amino]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
921	3-amino-6-(3-[[[3-bromo-5-methylphenyl]amino]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
922	3-amino-6-(3-[[[2-bromophenyl]amino]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
923	5-[3-(aminomethyl)phenyl]-3-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazine-2-amine
924	3-amino-6-(3-[[[4-(4-methylpiperazin-1-yl)phenyl]methyl]amino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
925	3-amino-6-(3-[[[2-(4-fluorophenyl)ethyl]amino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
926	3-amino-6-(3-[[[2-morpholin-4-ylethyl]amino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
927	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[2R]-pyrrolidin-2-ylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
928	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[2S]-pyrrolidin-2-ylamino]carbonyl]phenyl)pyrazine-2-carboxamide
929	3-amino-6-(3-[(cyclopentylamino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
930	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[4-[(trifluoromethyl)oxy]phenyl]amino]carbonyl]amino]methyl]phenyl)pyrazine-2-carboxamide
931	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[[4-(trifluoromethyl)phenyl]amino]carbonyl]amino]methyl]phenyl)pyrazine-2-carboxamide
932	3-amino-6-(3-[[[4-(methoxy)phenyl]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
933	3-amino-6-(3-[[[biphenyl-4-yl]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
934	3-amino-6-(3-[[[furan-2-yl]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
935	3-amino-6-[3-[[[2,3-dihydro-1,4-benzodioxin-6-ylamino]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
936	3-amino-6-(3-[[[2,3-dihydro-1-benzofuran-5-yl]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
937	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-6-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
938	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2-thienylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
939	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-2-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
940	N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl})pyrazin-2-yl)phenyl)methyl]-2-chlorobenzamide
941	N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl})pyrazin-2-yl)phenyl)methyl]-4-chlorobenzamide
942	N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl})pyrazin-2-yl)phenyl)methyl]-4-iodobenzamide
943	N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl})pyrazin-2-yl)phenyl)methyl]-3,5-difluorobenzamide
944	N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl})pyrazin-2-yl)phenyl)methyl]-4-bromo-2-fluorobenzamide
945	N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl})pyrazin-2-yl)phenyl)methyl]-4-bromo-2-chlorobenzamide
946	3-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]-5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazin-2-amine
947	3-amino-N-azepan-3-yl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
948	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2-(2-thienyl)-1,3-thiazol-4-yl)carbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
949	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((5-(2-thienyl)pyridin-3-yl)carbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
950	3-amino-6-(3-(((4-(1,1-dimethylethyl)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
951	3-amino-6-(3-(((2,4-dichloro-6-methylphenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
952	3-amino-6-(3-(((3,5-dichlorophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
953	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2-thienylsulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
954	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((3-thienylsulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
955	3-amino-6-(3-(((4-chlorophenyl)sulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
956	3-amino-6-(3-(((1,3-benzodioxol-5-ylamino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
957	3-amino-6-(3-(((4-(methyloxy)phenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
958	3-amino-6-(3-(((4-fluorophenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
959	3-amino-6-(3-([2-(diethylamino)ethyl]oxy)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
960	3-amino-6-[3-[(2-morpholin-4-ylethyl)oxy]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
961	3-amino-6-[3-(methyloxy)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
962	3-amino-6-[3-(ethyloxy)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
963	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-(1,2,3-thiadiazol-4-yl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
964	3-amino-6-[3-[(2-(ethylamino)ethyl)oxy]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
965	phenylmethyl 4-(2-[[3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl]pyrazin-2-yl)phenyl]oxy)ethyl)piperazine-1-carboxylate
966	3-amino-6-[3-[[[4-(dimethylamino)phenyl]amino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
967	3-amino-N-azepan-3-yl-6-[3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazine-2-carboxamide
968	3-amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
969	3-amino-6-[3-[[[(1S,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide
970	3-amino-N-[(3S)-azepan-3-yl]-6-[3-[[[(1S,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
971	3-amino-6-[3-[[[4-(chlorophenyl)amino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
972	3-amino-6-[3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide
973	3-amino-N-[(3S)-azepan-3-yl]-6-[3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazine-2-carboxamide
974	3-amino-N-[(3S)-azepan-3-yl]-6-(3-methylphenyl)pyrazine-2-carboxamide
975	3-amino-6-[3-[[2-(methylphenyl)acetyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
976	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[3-(trifluoromethyl)phenyl]acetyl]amino]phenyl]pyrazine-2-carboxamide
977	3-amino-6-[3-[[4-(methyloxy)phenyl]acetyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
978	3-amino-6-[3-[[6-(methylpyridin-3-yl)carbonyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
979	3-amino-6-[3-[[3-(fluorophenyl)acetyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
980	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[pyridin-3-ylacetyl]amino]phenyl]pyrazine-2-carboxamide
981	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[pyridin-3-ylcarbonyl]amino]phenyl]pyrazine-2-carboxamide
982	3-amino-6-[3-[[2,5-difluorophenyl]acetyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
983	3-amino-6-[3-(((4-fluorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
984	3-amino-6-[3-(((3-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
985	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
986	3-Amino-6-[3-(((5-(4-nitro-phenyl)-furan-2-carbonyl)-amino)-methyl)-phenyl]-pyrazine-2-carboxylic acid (3S)-piperidin-3-ylamide
987	3-amino-6-[3-(((3-hydroxypyridin-2-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
988	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-((trifluoromethyl)oxy)phenyl)oxy)acetyl)amino)methyl]phenyl]pyrazine-2-carboxamide
989	3-amino-6-[3-(((5-methylisoxazol-3-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
990	3-amino-6-[3-(((isoxazol-5-ylcarbonyl)amino)methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
991	3-amino-6-[3-(((4-methyl-1,2,3-thiadiazol-5-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
992	3-amino-6-[3-(((2,5-dichloro-3-thienyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
993	3-amino-6-[3-(((5-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
994	3-amino-6-[3-(((2-chloro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
995	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
996	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
997	3-amino-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
998	3-amino-6-[3-(((2-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
999	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1000	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1001	3-amino-6-[3-(((2-fluoro-4-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1002	3-amino-6-[3-(((2-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1003	3-amino-6-[3-(((2,3-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1004	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide

#	Name
1005	3-amino-6-[3-(((4-chloro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1006	3-amino-6-[3-(((3-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1007	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1008	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1009	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1010	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1011	3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1012	3-amino-6-[3-(((6-chloro-2-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1013	3-amino-6-[3-(((2-chloro-6-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1014	3-amino-6-[3-(((2-chloro-3,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1015	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1016	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,4-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1017	3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1018	3-amino-6-[3-(((3-fluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1019	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1020	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1021	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1022	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1023	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1024	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1025	3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1026	3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
1027	3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1028	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1029	3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1030	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1031	3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1032	3-amino-6-[3-(((3-bromo-5-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1033	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1034	3-amino-6-[3-(((biphenyl-3-yl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1035	3-amino-6-[3-(((2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1036	3-amino-6-[3-(((2,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1037	3-amino-6-[3-(((3,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1038	3-amino-6-[3-(((2,4-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1039	3-amino-6-[3-(((3-methoxyphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1040	3-amino-6-[3-(((phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1041	3-amino-6-[3-(((3,5-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1042	3-amino-6-[3-(((3,4-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1043	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide

[081] Another aspect of the invention is a pharmaceutical composition comprising a compound according to any one of paragraphs [0039]-[0080] and a pharmaceutically acceptable carrier.

[082] Another aspect of the invention is a metabolite of the compound or the pharmaceutical composition according to any one of paragraphs [0039]-[0081].

[083] Another aspect of the invention is a method of modulating the *in vivo* activity of a kinase, the method comprising administering to a subject an effective amount of the

compound or the pharmaceutical composition according to any of paragraphs [0039]-[0081].

[084] Another aspect of the invention is a method according to paragraph [0083], wherein the kinase is Chk1.

[085] Another aspect of the invention is a method according to paragraph [0084], wherein modulating the *in vivo* activity of the kinase comprises inhibition of said kinase.

[086] Another aspect of the invention is a method of treating diseases or disorders associated with uncontrolled, abnormal, and/or unwanted cellular activities, the method comprising administering a therapeutically effective amount of to a mammal in need thereof and the compound as described in any one of paragraphs [0039]-[0080].

[087] Another aspect of the invention is a method of inhibiting proliferation of cancerous cells, the method comprising contacting the cells or causing the cells to be contacted with a cancer therapeutic and the compound as described in any one of paragraphs [0039]-[0080].

[088] Another aspect of the invention is a method according to paragraph [0087], wherein the cancer therapeutic is a DNA damaging agent.

[089] Yet another aspect of the invention is a method of diminishing proliferation of cancerous cells in a patient, the method comprising administering to the patient the compound as described in any one of paragraphs [0039]-[0080] and one or more cancer therapeutic agents, preferably with a pharmaceutically acceptable carrier. Exemplary anti-cancer agents suitable for use (including analogs, pharmaceutically acceptable forms, such as salts and in liposomes) in this aspect of the invention are: 1. *Topoisomerase I inhibitors*: Camptothecin, Topotecan, 9-Nitrocamptothecin, 9-Aminocamptothecin, Karenitecin, Irinotecan, and the like; 2. *Topoisomerase II inhibitors*: Etoposide, Etoposide Phosphate, Teniposide, Amsacrine, Epipodophyllotoxin derivatives, Razoxane, Dexrazoxane (Zinecard), and the like; 3. *Classical Alkylating agents*: Nitrogen Mustards such as Mechlorethamine, Cyclophosphamide, Ifosfamide, Chlorambucil, Melphalan, and the like; Aziridines such as, Thiotepa, Trenimon, Triethylenemelamine, and the like; Epoxides such as Dianhydrogalactitol, Dibromodulcitol, and the like; Alkyl Alkane Sulfonates such as Busulfan, dimethylsulfate, and the like; Nitrosoureas such as Chloroethylnitrosourea, BCNU, CCNU (lomustine), Methyl-CCNU (semustine), Streptozotocin, Chlorozotocin, and the like; Alkylating agent-steroid conjugates such as

Prednimustine (chlorambucil-prednisolone), Estramustine (nornitrogen mustard-estradiol), and the like; 4. *Non-classical alkylating agents*: Procarbazine, Dacarbazine, Hexamethylmelamine, Pentamethylmelamine, Temozolomide, and the like; 5. *Other DNA damaging/binding agents*: Cisplatin such as Cisplatin, Carboplatin, Oxaliplatin, Bleomycin, and the like; Antibiotics such as Dactinomycin, Mithramycin, Mitomycin C, and the like; Anthracyclines/Anthracenediones (note - also considered topo II inhibitors) such as Daunorubicin, Doxorubicin, Epirubicin, Idarubicin, and the like; 6. *Antimetabolites*: Antifolates such as Methotrexate, Edatrexate, Trimethoprim, Nolatrexed, Raltitrexed (Tomudex), Hydroxyurea, and the like; Nucleic acid analogs such as 5-fluorouracil, Ftorafur, Capecitabine, Furtulon, Eniluracil, ara-C (Cytosine arabinose), 5-azacytidine, Gemcitabine, Mercaptopurine, Thioguanine, Pentostatin, and the like; 7. *Ribonucleic acid related*: Antisense DNA, antisense RNA, antisense DNA/RNA hybrids, ribozymes, and the like; 8. *Radiation*: ultraviolet and/or other; 9. *Vinca Alkaloids*: Vincristine, Vinblastine, and the like; 10. *Other anti-cancer agents*: those having mechanisms of action that may or may not involve DNA damage; 11. *Taxanes*: Paclitaxel, Docetaxel, and the like; 12. *Enzymes*: L-Asparaginase and the like; 13. *Natural Products*; 14. *Miscellaneous Agents*: kinase inhibitors, Imatinib; 15. Mitotane; 16. Aminoglutethimide; 17. *Hormones and Antagonists*: Diethylstilbestrol, Ethinyl estradiol, Tamoxifen, Anastrozole, Testosterone propionate, Fluoxymesterone, Flutamide, Leuprolide, Prednisone, Hydroxyprogesterone caproate, Medroxyprogesterone acetate, Megestrol acetate; 18. *Biological Response Modifiers*: Interferon-alfa, Interleukin, and the like.

[090] Other agents suitable for use in combination with a compound of the invention are disclosed in "Cancer Chemotherapy and Biotherapy: Principles and Practice," Third edition, B. A. Chabner and D. L. Longo, eds., 2001, Lippincott Williams and Wilkins, Philadelphia, U.S.A.; P. Calabresi and B. A. Chabner, "Chemotherapy of Neoplastic Diseases" in "Goodman and Gilman's The Pharmacological Basis of Therapeutics," Tenth edition, J. G. Hardman and L. E. Limbird, eds., 2001, McGraw-Hill, New York, USA, pp. 1381 - 1388; and B. A. Chabner, D. P. Ryan, L. Paz-Ares, R. Garcia-Carbonero, and P. Calabresi, "Antineoplastic Agents" in "Goodman and Gilman's The Pharmacological Basis of Therapeutics," Tenth edition, J. G. Hardman and L. E. Limbird, eds., 2001, McGraw-Hill, New York, USA, pp. 1389 - 1459.

[091] Another aspect of the invention is a method of enhancing the efficacy of a disease therapy that inflicts DNA damage, the method comprising combining the disease therapy with administration of a compound or pharmaceutical composition as described in any one of paragraphs [0023]-[0066]. As an example, photochemotherapy of dermatological conditions such as psoriasis employs ultraviolet radiation in combination with psoralen type photosensitizers to cause DNA damage. In one embodiment of this aspect of the invention, such photochemotherapy is combined with administration of a compound or pharmaceutical composition as described in any one of paragraphs [0039]-[0081].

Definitions

[092] As used in the present specification, the following words and phrases are generally intended to have the meanings as set forth below, except to the extent that the context in which they are used indicates otherwise or they are expressly defined to mean something different.

[093] "Alkyl" is intended to include linear, branched, or cyclic hydrocarbon structures and combinations thereof, inclusively. For example, "C₈ alkyl" may refer to an *n*-octyl, *iso*-octyl, cyclohexylethyl, and the like. Lower alkyl refers to alkyl groups of from one to eight carbon atoms. Examples of lower alkyl groups include methyl, ethyl, propyl, isopropyl, butyl, *s*-butyl, *t*-butyl, isobutyl, pentyl, cyclopentyl, hexyl, cyclohexyl, and the like. Higher alkyl refers to alkyl groups containing more than 6 carbon atoms. Exemplary alkyl groups are those of C₂₀ or below. Cycloalkyl is a subset of alkyl and includes cyclic hydrocarbon groups of from 3 to 13 carbon atoms. Examples of cycloalkyl groups include *c*-propyl, *c*-butyl, *c*-pentyl, norbornyl, adamantyl and the like. Cycloalkyl rings may have unsaturation but are not aromatic. In this application, alkyl refers to alkanyl, alkenyl, and alkynyl residues (and combinations thereof); it is intended to include cyclohexylmethyl, vinyl, allyl, isoprenyl, and the like. Thus when an alkyl residue having a specific number of carbons is named, all geometric isomers having that number of carbons are intended to be encompassed; thus either "butyl" or "C₄alkyl" is meant to include *n*-butyl, *sec*-butyl, isobutyl, *t*-butyl, isobutenyl and but-2-yne radicals, for example; "propyl" or "C₃alkyl" each include *n*-propyl, propenyl, and isopropyl, for example. Alkyls with variable numbers of carbons may be named by using number ranges as subscripts, as for example, lower alkyl is equivalent to C₁₋₈alkyl.

- [094] "Alkylene" refers to straight or branched chain divalent radical consisting solely of carbon and hydrogen atoms, containing no unsaturation and having from one to ten carbon atoms, e.g., methylene, ethylene, propylene, n-butylene and the like. Alkylene is a subset of alkyl, referring to the same residues as alkyl, but having two points of attachment and specifically fully saturated. Examples of alkylene include ethylene ($-\text{CH}_2\text{CH}_2-$), propylene ($-\text{CH}_2\text{CH}_2\text{CH}_2-$), 2-dimethylpropylene ($-\text{CH}_2\text{C}(\text{CH}_3)_2\text{CH}_2-$), and 2-cyclohexylpropylene ($-\text{CH}_2\text{CH}(\text{C}_6\text{H}_{11})\text{CH}_2-$).
- [095] "Alkylidene" refers to a straight or branched chain unsaturated divalent radical consisting solely of carbon and hydrogen atoms, having from two to ten carbon atoms, e.g., ethylidene, propylidene, n-butylidene, and the like. Alkylidene is a subset of alkyl, referring to the same residues as alkyl, but having two points of attachment and specifically double bond unsaturation. The unsaturation present includes at least one double bond and a double bond can exist between the first carbon of the chain and a carbon atom of the rest of the molecule to which it is attached.
- [096] "Alkylidyne" refers to a straight or branched chain unsaturated divalent radical consisting solely of carbon and hydrogen atoms having from two to ten carbon atoms, e.g., propylid-2-ynyl, n-butylid-1-ynyl, and the like. Alkylidyne is a subset of alkyl, referring to the same residues as alkyl, but having two points of attachment and specifically triple bond unsaturation. The unsaturation present includes at least one triple bond and a triple bond can exist between the first carbon of the chain and a carbon atom of the rest of the molecule to which it is attached.
- [097] Any of the above radicals, "alkylene," "alkylidene" and "alkylidyne," when optionally substituted, may contain alkyl substitution which itself contains unsaturation. For example, 2-(2-phenylethynyl-but-3-enyl)-naphthalene (IUPAC name) contains an n-butylid-3-ynyl radical with a vinyl substituent at the 2-position of said radical.
- [098] "Alkoxy" refers to the group -O-alkyl, for example including carbon atoms of a straight, branched, cyclic configuration, unsaturated chains, and combinations thereof attached to the parent structure through an oxygen atom. Examples include methoxy, ethoxy, propoxy, isopropoxy, cyclopropyloxy, cyclohexyloxy and the like. Lower-alkoxy refers to groups containing one to eight carbons.
- [099] "Substituted alkoxy" refers to the group -O-(substituted alkyl), the substitution on the alkyl group generally containing more than only carbon (as defined by alkoxy). One

exemplary substituted alkoxy group is "polyalkoxy" or -O- (optionally substituted alkylene)-(optionally substituted alkoxy), and includes groups such as -OCH₂CH₂OCH₃, and glycol ethers such as polyethyleneglycol and -O(CH₂CH₂O)_xCH₃, where x is an integer of between about 2 and about 20, in another example, between about 2 and about 10, and in a further example between about 2 and about 5. Another exemplary substituted alkoxy group is hydroxyalkoxy or -OCH₂(CH₂)_yOH, where y is for example an integer of between about 1 and about 10, in another example y is an integer of between about 1 and about 4. Thus, where a group is defined as -OR, where "R" is optionally substituted alkyl, then such a group would include, but not be limited to, hydroxyalkoxy, polyalkoxy, and the like.

[0100] "Alkoxy" refers to an alkoxy as a linking group, for example -OCH₂- and the like. Lower alkoxy then refers to groups containing one to eight carbons and an oxygen.

[0101] "Substituted alkoxy" refers to a linking group alkoxy, for example -OCH₂CH₂OCH₂-. Optionally substituted alkoxy refers to both alkoxy and substituted alkoxy.

[0102] "Acyl" refers to groups of from one to ten carbon atoms of a straight, branched, cyclic configuration, saturated, unsaturated and aromatic and combinations thereof, attached to the parent structure through a carbonyl functionality. One or more carbons in the acyl residue may be replaced by nitrogen, oxygen or sulfur as long as the point of attachment to the parent remains at the carbonyl. Examples include acetyl, benzoyl, propionyl, isobutyryl, t-butoxycarbonyl, benzyloxycarbonyl and the like. Lower-acyl refers to groups containing one to eight carbons.

[0103] "α-Amino Acids" refer to naturally occurring and commercially available amino acids and optical isomers thereof. Typical natural and commercially available α-amino acids are glycine, alanine, serine, homoserine, threonine, valine, norvaline, leucine, isoleucine, norleucine, aspartic acid, glutamic acid, lysine, ornithine, histidine, arginine, cysteine, homocysteine, methionine, phenylalanine, homophenylalanine, phenylglycine, ortho-tyrosine, meta-tyrosine, para-tyrosine, tryptophan, glutamine, asparagine, proline and hydroxyproline. A "side chain of an α-amino acid" refers to the radical found on the α-carbon of an α-amino acid as defined above, for example, hydrogen (for glycine), methyl (for alanine), benzyl (for phenylalanine), and the like.

- [0104] "Amino" refers to the group -NH₂. "Substituted amino," refers to the group -NHR or -NRR where each R is independently selected from the group: optionally substituted alkyl, optionally substituted alkoxy, optionally substituted aryl, optionally substituted heterocyclyl, acyl, carboxy, alkoxycarbonyl, sulfanyl, sulfinyl and sulfonyl, e.g., diethylamino, methylsulfonylamino, furanyl-oxy-sulfonamino.
- [0105] "Aryl" refers to aromatic 6- to 14-membered carbocyclic rings include, e.g., benzene, naphthalene, indane, tetralin, fluorene and the like.
- [0106] "Arylene" refers to an "aryl" that is a linker between two other groups. For example, for G-V-B, "phenylene" for V refers to a benzene ring system that links G and B, that is, G and B are each attached to different carbons of V, leaving four potential sites for further substitution on V.
- [0107] "Arylalkyl" refers to a residue in which an aryl moiety is attached to a parent structure via one of an alkylene, alkylidene, or alkylidyne radical. Examples include benzyl, phenethyl, phenylvinyl, phenylallyl and the like. The aryl, alkylene, alkylidene, or alkylidyne radical portion of an arylalkyl group may be optionally substituted. "Lower arylalkyl" refers to an arylalkyl where the "alkyl" portion of the group has one to eight carbons.
- [0108] "Halogen" or "halo" refers to fluorine, chlorine, bromine or iodine. Dihaloaryl, dihaloalkyl, trihaloaryl etc. refer to aryl and alkyl substituted with a plurality of halogens, but not necessarily a plurality of the same halogen; thus 4-chloro-3-fluorophenyl is within the scope of dihaloaryl.
- [0109] "Heteroatom" refers to O, S, N, or P.
- [0110] "Heterocyclyl" refers to a stable 3- to 15-membered ring radical that consists of carbon atoms and from one to five heteroatoms selected from the group consisting of nitrogen, phosphorus, oxygen and sulfur. For purposes of this invention, the heterocyclyl ring radical may be a monocyclic, bicyclic or tricyclic ring system, which may include fused or bridged ring systems, either aromatic, saturated, or combinations thereof; and the nitrogen, phosphorus, carbon or sulfur atoms in the heterocyclyl ring radical may be optionally oxidized to various oxidation states, for example for the purposes of this invention and to negate undo repetition in the description, the corresponding N-oxide of pyridine derivatives, and the like, are understood to be included as compounds of the invention. In addition, the nitrogen atom may be optionally quaternized; and the ring

radical may be partially or fully saturated or aromatic. Examples of such heterocyclyl ring radicals include, but are not limited to, azetidiny, acridiny, benzodioxoly, benzodioxany, benzofurany, carbazoy, cinnoliny, dioxolany, indoliziny, naphthyridiny, perhydroazepiny, phenaziny, phenothiaziny, phenoxaziny, phthalaziny, pteridiny, puriny, quinoxaliny, quinoxaliny, quinoliny, isoquinoliny, tetrazoy, tetrahydroisoquinolyl, piperidiny, piperaziny, 2-oxopiperaziny, 2-oxopiperidiny, 2-oxopyrrolidiny, 2-oxoazepiny, azepiny, pyrroly, 4-piperidony, pyrrolidiny, pyrazoly, pyrazolidiny, imidazoly, imidazoliny, imidazolidiny, dihydropyridiny, tetrahydropyridiny, pyridiny, pyraziny, pyrimidiny, pyridaziny, oxazoly, oxazoliny, oxazolidiny, triazoly, indany, isoxazoly, isoxazolidiny, morpholiny, thiazoly, thiazoliny, thiazolidiny, isothiazoly, quinuclidiny, isothiazolidiny, indoly, isoindoly, indoliny, isoindoliny, octahydroindoly, octahydroisoindoly, quinoly, isoquinoly, decahydroisoquinoly, benzimidazolyl, thiadiazoly, benzopyrany, benzothiazoly, benzoxazoly, furyl, tetrahydrofuryl, tetrahydropyrany, thienyl, benzothielyl, thiamorpholiny, thiamorpholiny sulfoxide, thiamorpholiny sulfone, dioxaphospholany, and oxadiazoly.

[0111] "Heteroalicyclic" refers specifically to a non-aromatic heterocyclyl ring system radical. Heteroalicyclic ring systems include saturated and partially bridged ring systems having geometries [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], [3.1.0], [3.3.3], [3.3.2], [3.3.1], [3.2.2], [3.2.1], [2.2.2], [2.2.1], and the like.

[0112] "Heteroaryl" refers specifically to an aromatic heterocyclyl ring system radical.

[0113] "Heteroarylene" refers to a "heteroaryl" that is a linker between two other groups. For example, for M-P-L, "pyridylene" for P refers to a pyridine ring system that links M and L, that is, M and L are each attached to different atoms of P, leaving four potential sites for further substitution on P.

[0114] "Heterocyclylalkyl" refers to a residue in which a heterocyclyl ring is attached to a parent structure via one of an alkylene, alkylidene, or alkylidyne radical. Examples include (4-methylpiperazin-1-yl) methyl, (morpholin-4-yl) methyl, 2-(oxazolin-2-yl) ethyl, 4-(4-methylpiperazin-1-yl)-2-butenyl, and the like. The heterocyclyl, alkylene, alkylidene, or alkylidyne radical portion of an arylalkyl group may be optionally substituted. "Lower heterocyclylalkyl" refers to an arylalkyl where the "alkyl" portion of the group has one to eight carbons.

[0115] The term “imino” refers to a substitution on a carbon atom, more specifically to a doubly bonded nitrogen. For example, an imine, an amidine, and an oxime, all contain the “imino” group.

[0116] “Optional” or “optionally” means that the subsequently described event or circumstance may or may not occur, and that the description includes instances where said event or circumstance occurs and instances in which it does not. It will be understood by those skilled in the art with respect to any group containing one or more substituents that such groups are not intended to introduce any substitution or substitution patterns (e.g., substituted alkyl includes optionally substituted cycloalkyl groups, which in turn are defined as including optionally substituted alkyl groups, potentially *ad infinitum*) that are sterically impractical and/or synthetically non-feasible. “Optionally substituted” refers to all subsequent modifiers in a term, for example in the term “optionally substituted C₁₋₈alkylaryl,” optional substitution may occur on both the “C₁₋₈alkyl” portion and the “aryl” portion of the molecule; and for example, optionally substituted alkyl includes optionally substituted cycloalkyl groups, which in turn are defined as including optionally substituted alkyl groups, potentially *ad infinitum*. If a heterocyclic ring is “optionally substituted,” then both the carbon and any heteroatoms in the ring may be substituted thereon. Examples of optional substitution include, but are not limited to alkyl, halogen, alkoxy, hydroxy, oxo, carbamyl, acylamino, sulfonamido, carboxy, alkoxycarbonyl, acyl, alkylthio, alkylsulfonyl, nitro, cyano, amino, alkylamino, cycloalkyl and the like. Thus, for example, if a group “-C(O)R” is described, where “R” is optionally substituted alkyl, then “R” would include, but not be limited to, -CH₂Ph, -CH₂CH₂OPh, -CH=CHPhCH₃, -C₃H₄CH₂N(H)Ph, and the like.

[0116] The term “ortho” is normally used in reference to relative position of two substituents on a benzene ring; however, in this application the term “ortho” is meant to apply to other aromatic ring systems where two substituents reside on adjacent carbons. For example, 3-bromo-4-fluoro-thiophene possesses a bromo group and a fluoro group which have an ortho, or 1,2-positional relationship, to each other.

[0117] The term “oxo” refers to a substitution on a carbon atom, more specifically to a doubly bonded oxygen. For example, an oxo-morpholine, a cyclohexanone, and an acyl group, all contain the “oxo” functionality.

[0118] “Saturated bridged ring system” refers to a bicyclic or polycyclic ring system that is not aromatic. Such a system may contain isolated or conjugated unsaturation, but not

aromatic or heteroaromatic rings in its core structure (but may have aromatic substitution thereon). For example, 2,3,3a,4,7,7a-hexahydro-1H-indene and 1,2,3,4,4a,5,8,8a-octahydro-naphthalene are included in the class "saturated bridged ring system."

[0119] "Substituted" alkyl, aryl, and heterocyclyl, refer respectively to alkyl, aryl, and heterocyclyl, wherein one or more (for example up to about 5, in another example, up to about 3) hydrogen atoms are replaced by a substituent independently selected from the group: optionally substituted alkyl (e.g., fluoroalkyl), optionally substituted alkoxy, alkylenedioxy (e.g. methylenedioxy), optionally substituted amino (e.g., alkylamino and dialkylamino), optionally substituted amidino, optionally substituted aryl (e.g., phenyl), optionally substituted arylalkyl (e.g., benzyl), optionally substituted aryloxy (e.g., phenoxy), optionally substituted arylalkyloxy (e.g., benzyloxy), carboxy (-COOH), carboalkoxy (i.e., acyloxy or -OOCR), carboxyalkyl (i.e., esters or -COOR), carboxamido, aminocarbonyl, benzyloxycarbonylamino (CBZ-amino), cyano, carbonyl, halogen, hydroxy, optionally substituted heterocyclylalkyl, optionally substituted heterocyclyl, nitro, sulfanyl, sulfinyl, sulfonyl, and thio.

[0120] "Sulfanyl" refers to the groups: -S-(optionally substituted alkyl), -S-(optionally substituted aryl), and -S-(optionally substituted heterocyclyl).

[0121] "Sulfinyl" refers to the groups: -S(O)-H, -S(O)-(optionally substituted alkyl), -S(O)-(optionally substituted aryl), and -S(O)-(optionally substituted heterocyclyl).

[0122] "Sulfonyl" refers to the groups: -S(O₂)-H, -S(O₂)-(optionally substituted alkyl), -S(O₂)-(optionally substituted aryl), -S(O₂)-(optionally substituted heterocyclyl), -S(O₂)-(optionally substituted alkoxy), -S(O₂)-(optionally substituted aryloxy), and -S(O₂)-(optionally substituted heterocycliloxy).

[0123] The term "thiono" refers to a substitution on a carbon atom, more specifically to a doubly bonded sulfur. For example, a thioketone and a thioamide both contain the "thiono" functionality.

[0124] "Yield" for each of the reactions described herein is expressed as a percentage of the theoretical yield.

[0125] In some embodiments, as will be appreciated by those in the art, two adjacent groups on an aromatic system may be fused together to form a ring structure. The fused ring structure may contain heteroatoms and may be optionally substituted with one or

more groups. It should additionally be noted that saturated carbons of such fused groups (i.e. saturated ring structures) may contain two substitution groups.

[0126] Some of the compounds of the invention may have imino, amino, oxo or hydroxy substituents off aromatic heterocyclyl ring systems. For purposes of this disclosure, it is understood that such imino, amino, oxo or hydroxy substituents may exist in their corresponding tautomeric form, i.e., amino, imino, hydroxy or oxo, respectively. As well, tautomeric forms of ring conjugation, for example in a triazole, are understood to be within the scope of the invention, when formulae depict only one such tautomer.

[0127] Compounds of the invention are generally named using ACD/Name (available from Advanced Chemistry Development, Inc. of Toronto, Canada). This software derives names from chemical structures according to systematic application of the nomenclature rules agreed upon by the International Union of Pure and Applied Chemistry (IUPAC), International Union of Biochemistry and Molecular Biology (IUBMB), and the Chemical Abstracts Service (CAS).



[0128] The compounds of the invention, or their pharmaceutically acceptable salts, may have asymmetric carbon atoms, oxidized sulfur atoms or quaternized nitrogen atoms in their structure.

[0129] The compounds of the invention and their pharmaceutically acceptable salts may exist as single stereoisomers, racemates, and as mixtures of enantiomers and diastereomers. The compounds may also exist as geometric isomers. All such single stereoisomers, racemates and mixtures thereof, and geometric isomers are intended to be within the scope of this invention.

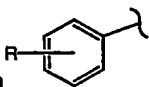
[0130] For linking groups, e.g. $-\text{OCH}_2-$ or $-\text{C}(=\text{O})\text{N}(\text{R})-$, it is understood that either of the linking groups two partners may be bound to either end of the linking group. Put another way, linking groups are understood to describe both possible orientations between the partners that they link. Therefore for A-B-C, where B is $-\text{OCH}_2-$, both bonding schemes A- OCH_2 -B and A- CH_2O -B are described. In the case of substituted linking groups, e.g. substituted alkoxyl, it is assumed that the substitution is in addition to the two groups to which the linking group is attached.

[0131] Methods for the preparation and/or separation and isolation of single stereoisomers from racemic mixtures or non-racemic mixtures of stereoisomers are well known in the art. For example, optically active (R)- and (S)- isomers may be prepared using chiral

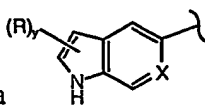
synthons or chiral reagents, or resolved using conventional techniques. When desired, the R- and S-isomers may be resolved by methods known to those skilled in the art, for example by: formation of diastereoisomeric salts or complexes which may be separated, for example, by crystallization; via formation of diastereoisomeric derivatives which may be separated, for example, by crystallization, gas-liquid or liquid chromatography; selective reaction of one enantiomer with an enantiomer-specific reagent, for example enzymatic oxidation or reduction, followed by separation of the modified and unmodified enantiomers; or gas-liquid or liquid chromatography in a chiral environment, for example on a chiral support, such as silica with a bound chiral ligand or in the presence of a chiral solvent. It will be appreciated that where a desired enantiomer is converted into another chemical entity by one of the separation procedures described above, a further step may be required to liberate the desired enantiomeric form. Alternatively, specific enantiomer may be synthesized by asymmetric synthesis using optically active reagents, substrates, catalysts or solvents, or by converting one enantiomer to the other by asymmetric transformation. For a mixture of enantiomers, enriched in a particular enantiomer, the major component enantiomer may be further enriched (with concomitant loss in yield) by recrystallization.

[0132] The symbol “-” means a single bond, “=” means a double bond, “≡” means a triple bond, and “” refers to a group on a double-bond as occupying either position on the terminus of a double bond to which the symbol is attached. When a group is depicted removed from its parent formula, the “” symbol will be used at the end of the bond which was theoretically cleaved in order to separate the group from its parent structural formula.

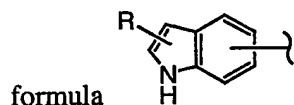
[0133] When a group “R” is depicted as existing on a ring system, as for example in the

formula , then a substituent “R” may reside on any atom of the ring system, assuming replacement of a depicted, implied, or expressly defined hydrogen from one of the ring atoms, so long as a stable structure is formed.

[0134] When a group “R” is depicted as existing on a fused ring system, as for example in

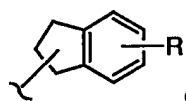
the formula , then a substituent “R” may reside on any atom of the fused ring system, assuming replacement of a depicted (e.g. the -NH- in the formula above),

implied (e.g. as in the formula above, where the hydrogens are not shown but understood to be present), or expressly defined hydrogen (e.g. where in the formula above, "X" equals =CH-) from one of the ring atoms, so long as a stable structure is formed. In the example depicted, the "R" group may reside on either the 5-membered or the 6-membered ring of the fused ring system. In the formula depicted above, when y is 2 for example, then the two "R's" may reside on any two atoms of the ring system, again assuming each replaces a depicted, implied, or expressly defined hydrogen on the ring. When there are more than one such depicted "floating" groups, as for example in the

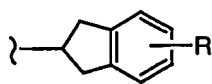


formula, where there are two groups, namely, the "R" and the bond indicating attachment to a parent structure. In such cases, the "floating" groups may reside on any atoms of the ring system, again assuming each floating group replaces a depicted, implied, or expressly defined unique hydrogen on the ring.

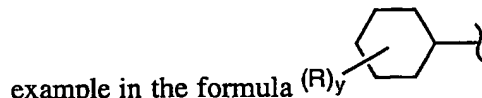
[0135] "Substituted cycloalkyl," in this application, is meant to include bridged or fused ring systems wherein an all-carbon bridge portion of such ring systems, along with the two bridgehead carbons, is considered as a cycloalkyl. For example, the formula



depicts a bridged ring system that has "R" substitution and is itself a substituent of a parent compound (as denoted by the "∼" symbol, *supra*). As mentioned above, either of the depicted "floating" groups may reside on either the alkyl bridge or the aromatic ring of the depicted bridged ring system. In this application, for such ring systems, when the parent compound is bonded to the alkyl bridge, as in



[0136] When a group "R" is depicted as existing on a saturated ring system, as for



example in the formula (R)_y, where "y" can be more than one, assuming each replaces a currently depicted, implied, or expressly defined hydrogen on the ring, then where the resulting structure is stable, two "R's" may reside on the same carbon. A simple example is when R is a methyl group, then in this instance there would exist a geminal dimethyl on a carbon of the depicted ring. In another example, two R's on the same carbon, including that carbon, may form a ring, thus creating a spirocyclic ring structure with the depicted ring.

[0137] "Patient" for the purposes of the present invention includes humans and other animals, particularly mammals, and other organisms. Thus the methods are applicable to both human therapy and veterinary applications. In a preferred embodiment the patient is a mammal, and in a most preferred embodiment the patient is human.

[0138] "Kinase-dependent diseases or conditions" refer to pathologic conditions that depend on the activity of one or more protein kinases. Kinases either directly or indirectly participate in the signal transduction pathways of a variety of cellular activities including proliferation, adhesion, migration, differentiation and invasion.

[0139] While not wishing to be bound to theory, phosphatases can also play a role in "kinase-dependent diseases or conditions" as cognates of kinases. That is, kinases phosphorylate and phosphatases dephosphorylate, for example protein substrates. Therefore compounds of the invention, while modulating kinase activity as described herein, may also modulate, either directly or indirectly, phosphatase activity. This additional modulation, if present, may be synergistic (or not) to activity of compounds of the invention toward a related or otherwise interdependent kinase or kinase family. Thus, "kinase-dependent diseases or conditions" are those diseases that should be mitigated via treatment of a patient with compounds and/or formulations of the invention. Kinase-dependent diseases or conditions include diseases or disorders associated with uncontrolled, abnormal and/or unwanted cellular activities such as those set forth in paragraphs [0031]-[0038].

[0140] "Therapeutically effective amount" is an amount of a compound of the invention, that when administered to a patient, ameliorates a symptom of the disease. The amount of a compound of the invention which constitutes a "therapeutically effective amount" will vary depending on the compound, the disease state and its severity, the age of the patient to be treated, and the like. The therapeutically effective amount can be determined routinely by one of ordinary skill in the art having regard to his own knowledge and to this disclosure.

[0141] "Cancer" refers to cellular-proliferative disease states, including but not limited to: Cardiac: sarcoma (angiosarcoma, fibrosarcoma, rhabdomyosarcoma, liposarcoma), myxoma, rhabdomyoma, fibroma, lipoma and teratoma; Lung: bronchogenic carcinoma (squamous cell, undifferentiated small cell, undifferentiated large cell, adenocarcinoma), alveolar (bronchiolar) carcinoma, bronchial adenoma, sarcoma, lymphoma, chondromatous hamartoma, mesothelioma; Gastrointestinal: esophagus (squamous cell

carcinoma, adenocarcinoma, leiomyosarcoma, lymphoma), stomach (carcinoma, lymphoma, leiomyosarcoma), pancreas (ductal adenocarcinoma, insulinoma, glucagonoma, gastrinoma, carcinoid tumors, vipoma), small bowel (adenocarcinoma, lymphoma, carcinoid tumors, Kaposi's sarcoma, leiomyoma, hemangioma, lipoma, neurofibroma, fibroma), large bowel (adenocarcinoma, tubular adenoma, villous adenoma, hamartoma, leiomyoma); Genitourinary tract: kidney (adenocarcinoma, Wilm's tumor [nephroblastoma], lymphoma, leukemia), bladder and urethra (squamous cell carcinoma, transitional cell carcinoma, adenocarcinoma), prostate (adenocarcinoma, sarcoma), testis (seminoma, teratoma, embryonal carcinoma, teratocarcinoma, choriocarcinoma, sarcoma, interstitial cell carcinoma, fibroma, fibroadenoma, adenomatoid tumors, lipoma); Liver: hepatoma (hepatocellular carcinoma), cholangiocarcinoma, hepatoblastoma, angiosarcoma, hepatocellular adenoma, hemangioma; Bone: osteogenic sarcoma (osteosarcoma), fibrosarcoma, malignant fibrous histiocytoma, chondrosarcoma, Ewing's sarcoma, malignant lymphoma (reticulum cell sarcoma), multiple myeloma, malignant giant cell tumor chordoma, osteochondroma (osteochondrogenous exostoses), benign chondroma, chondroblastoma, chondromyxofibroma, osteoid osteoma and giant cell tumors; Nervous system: skull (osteoma, hemangioma, granuloma, xanthoma, osteitis deformans), meninges (meningioma, meningiosarcoma, gliomatosis), brain (astrocytoma, medulloblastoma, glioma, ependymoma, germinoma [pinealoma], glioblastoma multiforme, oligodendroglioma, schwannoma, retinoblastoma, congenital tumors), spinal cord neurofibroma, meningioma, glioma, sarcoma); Gynecological: uterus (endometrial carcinoma), cervix (cervical carcinoma, pre-tumor cervical dysplasia), ovaries (ovarian carcinoma [serous cystadenocarcinoma, mucinous cystadenocarcinoma, unclassified carcinoma], granulosa-thecal cell tumors, Sertoli-Leydig cell tumors, dysgerminoma, malignant teratoma), vulva (squamous cell carcinoma, intraepithelial carcinoma, adenocarcinoma, fibrosarcoma, melanoma), vagina (clear cell carcinoma, squamous cell carcinoma, botryoid sarcoma (embryonal rhabdomyosarcoma), fallopian tubes (carcinoma); Hematologic: blood (myeloid leukemia [acute and chronic], acute lymphoblastic leukemia, chronic lymphocytic leukemia, myeloproliferative diseases, multiple myeloma, myelodysplastic syndrome), Hodgkin's disease, non-Hodgkin's lymphoma [malignant lymphoma]; Skin: malignant melanoma, basal cell carcinoma, squamous cell carcinoma, Kaposi's sarcoma, moles dysplastic nevi, lipoma, angioma, dermatofibroma, keloids, psoriasis; and Adrenal glands: neuroblastoma. Thus, the term

“cancerous cell” as provided herein, includes a cell afflicted by any one of the above-identified conditions.

[0142] “Pharmaceutically acceptable acid addition salt” refers to those salts that retain the biological effectiveness of the free bases and that are not biologically or otherwise undesirable, formed with inorganic acids such as hydrochloric acid, hydrobromic acid, sulfuric acid, nitric acid, phosphoric acid, and the like, as well as organic acids such as acetic acid, trifluoroacetic acid, propionic acid, glycolic acid, pyruvic acid, oxalic acid, maleic acid, malonic acid, succinic acid, fumaric acid, tartaric acid, citric acid, benzoic acid, cinnamic acid, mandelic acid, methanesulfonic acid, ethanesulfonic acid, p-toluenesulfonic acid, salicylic acid and the like.

[0143] “Pharmaceutically acceptable base addition salts” include those derived from inorganic bases such as sodium, potassium, lithium, ammonium, calcium, magnesium, iron, zinc, copper, manganese, aluminum salts and the like. Exemplary salts are the ammonium, potassium, sodium, calcium, and magnesium salts. Salts derived from pharmaceutically acceptable organic non-toxic bases include, but are not limited to, salts of primary, secondary, and tertiary amines, substituted amines including naturally occurring substituted amines, cyclic amines and basic ion exchange resins, such as isopropylamine, trimethylamine, diethylamine, triethylamine, tripropylamine, ethanolamine, 2-dimethylaminoethanol, 2-diethylaminoethanol, dicyclohexylamine, lysine, arginine, histidine, caffeine, procaine, hydrabamine, choline, betaine, ethylenediamine, glucosamine, methylglucamine, theobromine, purines, piperazine, piperidine, N-ethylpiperidine, polyamine resins, and the like. Exemplary organic bases are isopropylamine, diethylamine, ethanolamine, trimethylamine, dicyclohexylamine, choline, and caffeine. (See, for example, S. M. Berge, et al., “Pharmaceutical Salts,” J. Pharm. Sci., 1977;66:1-19 which is incorporated herein by reference.)

[0144] “Prodrug” refers to compounds that are transformed (typically rapidly) *in vivo* to yield the parent compound of the above formulae, for example, by hydrolysis in blood. Common examples include, but are not limited to, ester and amide forms of a compound having an active form bearing a carboxylic acid moiety. Examples of pharmaceutically acceptable esters of the compounds of this invention include, but are not limited to, alkyl esters (for example with between about 1 and about 6 carbons) wherein the alkyl group is a straight or branched chain. Acceptable esters also include cycloalkyl esters and arylalkyl esters such as, but not limited to benzyl. Examples of pharmaceutically

acceptable amides of the compounds of this invention include, but are not limited to, primary amides, and secondary and tertiary alkyl amides (for example with between about 1 and about 6 carbons). Amides and esters of the compounds of the present invention may be prepared according to conventional methods. Another example of a prodrug is a pyridine group; in many cases the pyridine group is oxidized to its corresponding N-oxide that may be a biologically active compound of the invention. Thus the "pyridine form" may be thought of as a prodrug. A thorough discussion of prodrugs is provided in T. Higuchi and V. Stella, "Pro-drugs as Novel Delivery Systems," Vol 14 of the A.C.S. Symposium Series, and in Bioreversible Carriers in Drug Design, ed. Edward B. Roche, American Pharmaceutical Association and Pergamon Press, 1987, both of which are incorporated herein by reference.

[0145] "Metabolite" refers to the break-down or end product of a compound or its salt produced by metabolism or biotransformation in the animal or human body; e.g., biotransformation to a more polar molecule such as by oxidation, reduction, or hydrolysis, or to a conjugate (see Goodman and Gilman, "The Pharmacological Basis of Therapeutics" 8^{sup}.th Ed., Pergamon Press, Gilman et al. (eds), 1990 for a discussion of biotransformation). As used herein, the metabolite of a compound of the invention or its salt may be the biologically active form of the compound in the body. In one example, a prodrug may be used such that the biologically active form, a metabolite, is released *in vivo*. In another example, a biologically active metabolite is discovered serendipitously, that is, no prodrug design *per se* was undertaken. An assay for activity of a metabolite of a compound of the present invention is known to one of skill in the art in light of the present disclosure.

[0146] In addition, the compounds of the present invention can exist in unsolvated as well as solvated forms with pharmaceutically acceptable solvents such as water, ethanol, and the like. In general, the solvated forms are considered equivalent to the unsolvated forms for the purposes of the present invention.

[0147] In addition, it is intended that the present invention cover compounds made either using standard organic synthetic techniques, including combinatorial chemistry or by biological methods, such as bacterial digestion, metabolism, enzymatic conversion, and the like.

[0148] "Treating" or "treatment" as used herein covers the treatment of a disease-state in a human, which disease-state is characterized by uncontrolled, abnormal or unwanted

cellular proliferation, differentiation, programmed cell death migration or chemoinvasion and invasion and includes at least one of: (i) preventing the disease-state from occurring in a human, in particular, when such human is predisposed to the disease-state but has not yet been diagnosed as having it; (ii) inhibiting the disease-state, i.e., arresting its development; and (iii) relieving the disease-state, i.e., causing regression of the disease-state. As is known in the art, adjustments for systemic versus localized delivery, age, body weight, general health, sex, diet, time of administration, drug interaction and the severity of the condition may be necessary, and will be ascertainable with routine experimentation by those skilled in the art.

General Administration

[0149] Administration of the compounds of the invention, or their pharmaceutically acceptable salts, in pure form or in an appropriate pharmaceutical composition, can be carried out via any of the accepted modes of administration or agents for serving similar utilities. Thus, administration can be, for example, orally, nasally, parenterally (intravenous, intramuscular, or subcutaneous), topically, transdermally, intravaginally, intravesically, intracisternally, or rectally, in the form of solid, semi-solid, lyophilized powder, or liquid dosage forms, such as for example, tablets, suppositories, pills, soft elastic and hard gelatin capsules, powders, solutions, suspensions, or aerosols, or the like, preferably in unit dosage forms suitable for simple administration of precise dosages.

[0150] The compositions will include a conventional pharmaceutical carrier or excipient and a compound of the invention as the/an active agent, and, in addition, may include other medicinal agents, pharmaceutical agents, carriers, adjuvants, etc. Compositions of the invention may be used in combination with anticancer or other agents that are generally administered to a patient being treated for cancer. Adjuvants include preserving, wetting, suspending, sweetening, flavoring, perfuming, emulsifying, and dispensing agents. Prevention of the action of microorganisms can be ensured by various antibacterial and antifungal agents, for example, parabens, chlorobutanol, phenol, sorbic acid, and the like. It may also be desirable to include isotonic agents, for example sugars, sodium chloride, and the like. Prolonged absorption of the injectable pharmaceutical form can be brought about by the use of agents delaying absorption, for example, aluminum monostearate and gelatin.

[0151] If desired, a pharmaceutical composition of the invention may also contain minor amounts of auxiliary substances such as wetting or emulsifying agents, pH buffering agents, antioxidants, and the like, such as, for example, citric acid, sorbitan monolaurate, triethanolamine oleate, butylated hydroxytoluene, etc.

[0152] Compositions suitable for parenteral injection may comprise physiologically acceptable sterile aqueous or nonaqueous solutions, dispersions, suspensions or emulsions, and sterile powders for reconstitution into sterile injectable solutions or dispersions. Examples of suitable aqueous and nonaqueous carriers, diluents, solvents or vehicles include water, ethanol, polyols (propyleneglycol, polyethyleneglycol, glycerol, and the like), suitable mixtures thereof, vegetable oils (such as olive oil) and injectable organic esters such as ethyl oleate. Proper fluidity can be maintained, for example, by the use of a coating such as lecithin, by the maintenance of the required particle size in the case of dispersions and by the use of surfactants.

[0153] One preferable route of administration is oral, using a convenient daily dosage regimen that can be adjusted according to the degree of severity of the disease-state to be treated.

[0154] Solid dosage forms for oral administration include capsules, tablets, pills, powders, and granules. In such solid dosage forms, the active compound is admixed with at least one inert customary excipient (or carrier) such as sodium citrate or dicalcium phosphate or (a) fillers or extenders, as for example, starches, lactose, sucrose, glucose, mannitol, and silicic acid, (b) binders, as for example, cellulose derivatives, starch, alginates, gelatin, polyvinylpyrrolidone, sucrose, and gum acacia, (c) humectants, as for example, glycerol, (d) disintegrating agents, as for example, agar-agar, calcium carbonate, potato or tapioca starch, alginic acid, croscarmellose sodium, complex silicates, and sodium carbonate, (e) solution retarders, as for example paraffin, (f) absorption accelerators, as for example, quaternary ammonium compounds, (g) wetting agents, as for example, cetyl alcohol, and glycerol monostearate, magnesium stearate and the like (h) adsorbents, as for example, kaolin and bentonite, and (i) lubricants, as for example, talc, calcium stearate, magnesium stearate, solid polyethylene glycols, sodium lauryl sulfate, or mixtures thereof. In the case of capsules, tablets, and pills, the dosage forms may also comprise buffering agents.

[0155] Solid dosage forms as described above can be prepared with coatings and shells, such as enteric coatings and others well known in the art. They may contain pacifying

agents, and can also be of such composition that they release the active compound or compounds in a certain part of the intestinal tract in a delayed manner. Examples of embedded compositions that can be used are polymeric substances and waxes. The active compounds can also be in microencapsulated form, if appropriate, with one or more of the above-mentioned excipients.

[0156] Liquid dosage forms for oral administration include pharmaceutically acceptable emulsions, solutions, suspensions, syrups, and elixirs. Such dosage forms are prepared, for example, by dissolving, dispersing, etc., a compound(s) of the invention, or a pharmaceutically acceptable salt thereof, and optional pharmaceutical adjuvants in a carrier, such as, for example, water, saline, aqueous dextrose, glycerol, ethanol and the like; solubilizing agents and emulsifiers, as for example, ethyl alcohol, isopropyl alcohol, ethyl carbonate, ethyl acetate, benzyl alcohol, benzyl benzoate, propyleneglycol, 1,3-butyleneglycol, dimethylformamide; oils, in particular, cottonseed oil, groundnut oil, corn germ oil, olive oil, castor oil and sesame oil, glycerol, tetrahydrofurfuryl alcohol, polyethyleneglycols and fatty acid esters of sorbitan; or mixtures of these substances, and the like, to thereby form a solution or suspension.

[0157] Suspensions, in addition to the active compounds, may contain suspending agents, as for example, ethoxylated isostearyl alcohols, polyoxyethylene sorbitol and sorbitan esters, microcrystalline cellulose, aluminum metahydroxide, bentonite, agar-agar and tragacanth, or mixtures of these substances, and the like.

[0158] Compositions for rectal administrations are, for example, suppositories that can be prepared by mixing the compounds of the present invention with for example suitable non-irritating excipients or carriers such as cocoa butter, polyethyleneglycol or a suppository wax, which are solid at ordinary temperatures but liquid at body temperature and therefore, melt while in a suitable body cavity and release the active component therein.

[0159] Dosage forms for topical administration of a compound of this invention include ointments, powders, sprays, and inhalants. The active component is admixed under sterile conditions with a physiologically acceptable carrier and any preservatives, buffers, or propellants as may be required. Ophthalmic formulations, eye ointments, powders, and solutions are also contemplated as being within the scope of this invention.

[0160] Generally, depending on the intended mode of administration, the pharmaceutically acceptable compositions will contain about 1% to about 99% by weight of a compound(s) of the invention, or a pharmaceutically acceptable salt thereof, and 99% to 1% by weight of a suitable pharmaceutical excipient. In one example, the composition will be between about 5% and about 75% by weight of a compound(s) of the invention, or a pharmaceutically acceptable salt thereof, with the rest being suitable pharmaceutical excipients.

[0161] Actual methods of preparing such dosage forms are known, or will be apparent, to those skilled in this art; for example, see Remington's Pharmaceutical Sciences, 18th Ed., (Mack Publishing Company, Easton, Pa., 1990). The composition to be administered will, in any event, contain a therapeutically effective amount of a compound of the invention, or a pharmaceutically acceptable salt thereof, for treatment of a disease-state in accordance with the teachings of this invention.

[0162] The compounds of the invention, or their pharmaceutically acceptable salts, are administered in a therapeutically effective amount which will vary depending upon a variety of factors including the activity of the specific compound employed, the metabolic stability and length of action of the compound, the age, body weight, general health, sex, diet, mode and time of administration, rate of excretion, drug combination, the severity of the particular disease-states, and the host undergoing therapy. The compounds of the present invention can be administered to a patient at dosage levels in the range of about 0.1 to about 1,000 mg per day. For a normal human adult having a body weight of about 70 kilograms, a dosage in the range of about 0.01 to about 100 mg per kilogram of body weight per day is an example. The specific dosage used, however, can vary. For example, the dosage can depend on a number of factors including the requirements of the patient, the severity of the condition being treated, and the pharmacological activity of the compound being used. The determination of optimum dosages for a particular patient is well known to those skilled in the art.

Utility of compounds of the invention as screening agents

[0163] To employ the compounds of the invention in a method of screening for candidate agents that bind to, for example a Chk1 receptor kinase, the protein is bound to a support, and a compound of the invention is added to the assay. Alternatively, the compound of

the invention is bound to the support and the protein is added. Classes of candidate agents among which novel binding agents may be sought include specific antibodies, non-natural binding agents identified in screens of chemical libraries, peptide analogs, etc. Of particular interest are screening assays for candidate agents that have a low toxicity for human cells. A wide variety of assays may be used for this purpose, including labeled *in vitro* protein-protein binding assays, electrophoretic mobility shift assays, immunoassays for protein binding, functional assays (phosphorylation assays, etc.) and the like.

[0164] The determination of the binding of the candidate agent to, for example, a Chk1 protein may be done in a number of ways. In one example, the candidate agent (the compound of the invention) is labeled, for example, with a fluorescent or radioactive moiety and binding determined directly. For example, this may be done by attaching all or a portion of the Chk1 protein to a solid support, adding a labeled agent (for example a compound of the invention in which at least one atom has been replaced by a detectable isotope), washing off excess reagent, and determining whether the amount of the label is that present on the solid support. Various blocking and washing steps may be utilized as is known in the art.

[0165] By "labeled" herein is meant that the compound is either directly or indirectly labeled with a label which provides a detectable signal, e.g., radioisotope, fluorescent tag, enzyme, antibodies, particles such as magnetic particles, chemiluminescent tag, or specific binding molecules, etc. Specific binding molecules include pairs, such as biotin and streptavidin, digoxin and antidigoxin etc. For the specific binding members, the complementary member would normally be labeled with a molecule which provides for detection, in accordance with known procedures, as outlined above. The label can directly or indirectly provide a detectable signal.

[0166] In some embodiments, only one of the components is labeled. For example, a Chk1 protein may be labeled at tyrosine positions using ^{125}I , or with fluorophores. Alternatively, more than one component may be labeled with different labels; using ^{125}I for the proteins, for example, and a fluorophor for the candidate agents.

[0167] The compounds of the invention may also be used as competitors to screen for additional drug candidates. "Candidate bioactive agent" or "drug candidate" or grammatical equivalents as used herein describe any molecule, e.g., protein, oligopeptide, small organic molecule, polysaccharide, polynucleotide, etc., to be tested for bioactivity.

They may be capable of directly or indirectly altering the cellular proliferation phenotype or the expression of a cellular proliferation sequence, including both nucleic acid sequences and protein sequences. In other cases, alteration of cellular proliferation protein binding and/or activity is screened. In the case where protein binding or activity is screened, some embodiments exclude molecules already known to bind to that particular protein. Exemplary embodiments of assays described herein include candidate agents, which do not bind the target protein in its endogenous native state, termed herein as "exogenous" agents. In one example, exogenous agents further exclude antibodies to Chk1.

[0168] Candidate agents can encompass numerous chemical classes, though typically they are organic molecules having a molecular weight of more than about 100 and less than about 2,500 daltons. Candidate agents comprise functional groups necessary for structural interaction with proteins, particularly hydrogen bonding and lipophilic binding, and typically include at least an amine, carbonyl, hydroxyl, ether, or carboxyl group, for example at least two of the functional chemical groups. The candidate agents often comprise cyclical carbon or heterocyclyl structures and/or aromatic or polyaromatic structures substituted with one or more of the above functional groups. Candidate agents are also found among biomolecules including peptides, saccharides, fatty acids, steroids, purines, pyrimidines, derivatives, structural analogs, or combinations thereof.

[0169] Candidate agents are obtained from a wide variety of sources including libraries of synthetic or natural compounds. For example, numerous means are available for random and directed synthesis of a wide variety of organic compounds and biomolecules, including expression of randomized oligonucleotides. Alternatively, libraries of natural compounds in the form of bacterial, fungal, plant and animal extracts are available or readily produced. Additionally, natural or synthetically produced libraries and compounds are readily modified through conventional chemical, physical and biochemical means. Known pharmacological agents may be subjected to directed or random chemical modifications, such as acylation, alkylation, esterification, amidification to produce structural analogs.

[0170] In one example, the binding of the candidate agent is determined through the use of competitive binding assays. In this example, the competitor is a binding moiety known to bind to Chk1, such as an antibody, peptide, binding partner, ligand, etc. Under certain

circumstances, there may be competitive binding as between the candidate agent and the binding moiety, with the binding moiety displacing the candidate agent.

[0171] In some embodiments, the candidate agent is labeled. Either the candidate agent, or the competitor, or both, is added first to Chk1 for a time sufficient to allow binding, if present. Incubations may be performed at any temperature that facilitates optimal activity, typically between 4°C and 40°C.

[0172] Incubation periods are selected for optimum activity, but may also be optimized to facilitate rapid high throughput screening. Typically between 0.1 and 1 hour will be sufficient. Excess reagent is generally removed or washed away. The second component is then added, and the presence or absence of the labeled component is followed, to indicate binding.

[0173] In one example, the competitor is added first, followed by the candidate agent. Displacement of the competitor is an indication the candidate agent is binding to Chk1 and thus is capable of binding to, and potentially modulating, the activity of the Chk1. In this embodiment, either component can be labeled. Thus, for example, if the competitor is labeled, the presence of label in the wash solution indicates displacement by the agent. Alternatively, if the candidate agent is labeled, the presence of the label on the support indicates displacement.

[0174] In an alternative embodiment, the candidate agent is added first, with incubation and washing, followed by the competitor. The absence of binding by the competitor may indicate the candidate agent is bound to Chk1 with a higher affinity. Thus, if the candidate agent is labeled, the presence of the label on the support, coupled with a lack of competitor binding, may indicate the candidate agent is capable of binding to Chk1.

[0175] It may be of value to identify the binding site of Chk1. This can be done in a variety of ways. In one embodiment, once Chk1 has been identified as binding to the candidate agent, the Chk1 is fragmented or modified and the assays repeated to identify the necessary components for binding.

[0176] Modulation is tested by screening for candidate agents capable of modulating the activity of Chk1 comprising the steps of combining a candidate agent with Chk1, as above, and determining an alteration in the biological activity of the Chk1. Thus, in this embodiment, the candidate agent should both bind to (although this may not be necessary), and alter its biological or biochemical activity as defined herein. The

methods include both *in vitro* screening methods and *in vivo* screening of cells for alterations in cell viability, morphology, and the like.

[0177] Alternatively, differential screening may be used to identify drug candidates that bind to native Chk1, but cannot bind to modified Chk1.

[0178] Positive controls and negative controls may be used in the assays. For example, all control and test samples are performed in at least triplicate to obtain statistically significant results. Incubation of samples is for a time sufficient for the binding of the agent to the protein. Following incubation, samples are washed free of non-specifically bound material and the amount of bound, generally labeled agent determined. For example, where a radiolabel is employed, the samples may be counted in a scintillation counter to determine the amount of bound compound.

[0179] A variety of other reagents may be included in the screening assays. These include reagents like salts, neutral proteins, e.g., albumin, detergents, etc which may be used to facilitate optimal protein-protein binding and/or reduce non-specific or background interactions. Also reagents that otherwise improve the efficiency of the assay, such as protease inhibitors, nuclease inhibitors, anti-microbial agents, etc., may be used. The mixture of components may be added in any order that provides for the requisite binding.

Abbreviations and their Definitions

[0180] The following abbreviations and terms have the indicated meanings throughout:

Ac	=	acetyl
ATP	=	adenosine triphosphate
BNB	=	4-bromomethyl-3-nitrobenzoic acid
Boc	=	t-butyloxy carbonyl
br	=	broad
Bu	=	butyl
C	=	degrees Celsius
c-	=	cyclo
CBZ	=	carbobenzoxy = benzyloxycarbonyl
d	=	doublet
dd	=	doublet of doublet
dt	=	doublet of triplet
DBU	=	diazabicyclo[5.4.0]undec-7-ene

DCM	=	dichloromethane = methylene chloride = CH_2Cl_2
DCE	=	dichloroethylene
DEAD	=	diethyl azodicarboxylate
DIC	=	diisopropylcarbodiimide
DIEA	=	N,N-diisopropylethyl amine
DMAP	=	4-N,N-dimethylaminopyridine
DMF	=	N,N-dimethylformamide
DMSO	=	dimethyl sulfoxide
DVB	=	1,4-divinylbenzene
EEDQ	=	2-ethoxy-1-ethoxycarbonyl-1,2-dihydroquinoline
EI	=	Electron Impact ionization
Et	=	ethyl
Fmoc	=	9-fluorenylmethoxycarbonyl
g	=	gram(s)
GC	=	gas chromatography
h or hr	=	hour(s)
HATU	=	0-(7-Azabenzotriazol-1-yl)-1,1,3,3-tetramethyluronium hexafluorophosphate
HMDS	=	hexamethyldisilazane
HOAc	=	acetic acid
HOBt	=	hydroxybenzotriazole
HPLC	=	high pressure liquid chromatography
L	=	liter(s)
M	=	molar or molarity
m	=	multiplet
Me	=	methyl
mesyl	=	methanesulfonyl
mg	=	milligram(s)
MHz	=	megahertz (frequency)
min	=	minute(s)
mL	=	milliliter(s)
mM	=	millimolar
mmol	=	millimole(s)
mol	=	mole(s)

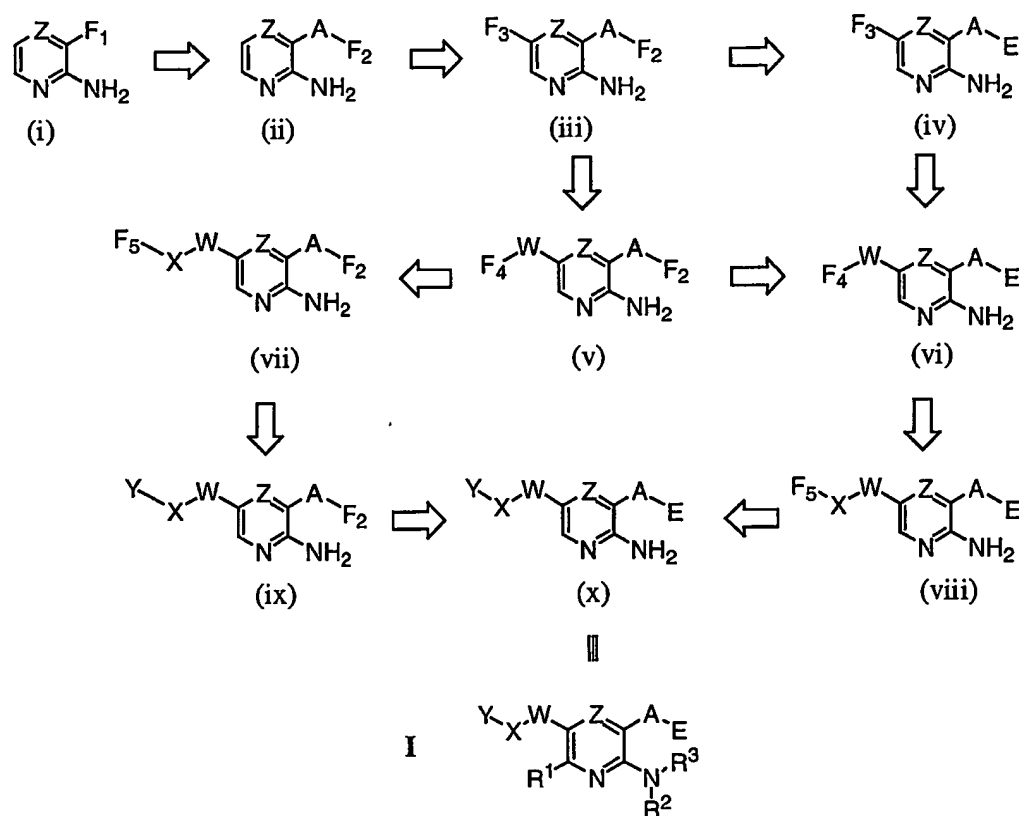
MS	=	mass spectral analysis
MTBE	=	methyl t-butyl ether
N	=	normal or normality
NBS	=	N-bromosuccinimide
NCS	=	N-chlorosuccinimide
nM	=	nanomolar
NMO	=	N-methylmorpholine oxide
NMR	=	nuclear magnetic resonance spectroscopy
PEG	=	polyethylene glycol
pEY	=	poly-glutamine, tyrosine
Ph	=	phenyl
PhOH	=	phenol
PfP	=	pentafluorophenol
PfPy	=	pentafluoropyridine
PPTS	=	pyridinium p-toluenesulfonate
Py	=	pyridine
PyBroP	=	bromo-tris-pyrrolidino-phosphonium hexafluorophosphate
q	=	quartet
RT	=	room temperature
Sat'd	=	saturated
s	=	singlet
s-	=	secondary
t-	=	tertiary
t or tr	=	triplet
TBDMS	=	t-butyldimethylsilyl
TES	=	triethylsilane
TFA	=	trifluoroacetic acid
THF	=	tetrahydrofuran
TMOF	=	trimethyl orthoformate
TMS	=	trimethylsilyl
tosyl	=	p-toluenesulfonyl
Trt	=	triphenylmethyl
uL	=	microliter(s)
uM	=	micromole(s) or micromolar

Synthesis of Compounds

- [0181] Scheme 1 depicts general synthetic routes for compounds of the invention and is not intended to be limiting. Specific examples are described subsequently to these general synthetic descriptions. In the generalizations below, specific reaction conditions, for example, reagents such as added bases, acids, solvents, temperature, and the like are not described so as not to confuse the discussion.
- [0182] The general routes, in conjunction with the specific examples, contain sufficient information to allow one skilled in the art to synthesize compounds of the invention. In Scheme 1, some substituents are not defined described as taking part in the synthesis of compounds of the invention. This is done purely for simplification of description of syntheses. Such substituents in formula I may be appended to the scaffold of formula I at any time during synthesis or may pre-exist on intermediates or starting materials used to make compounds of the invention, as would be understood by one skilled in the art. More specific examples are presented below to more fully describe the invention.
- [0183] Scheme 1 depicts examples of how compounds of the invention, consistent with formula I, are made (in these examples for simplicity of discussion, substituents R^1 , R^2 , and R^3 are -H, but this is not always the case). Generally, but not necessarily, a starting material will take the form of, for example, (i) or (ii). For example, (i) may take the form of a commercially available 3-amino-pyrazine (where Z is N). The group "F₁" represents functionality that can be converted to, or functionality that exists in, compounds of formula I. For example if F₁ represents a carboxylic acid group, then where A, of formula I, is a carbonyl, then (ii) may more appropriately represent the starting material. In other cases, A is introduced into (i) to make (ii). For example, if F₁ is a cyano group, then in some instances the cyano group in (i) is converted to a triazole ring system to produce A in (ii).
- [0184] F₂ represents a functionality that is ultimately converted to, or used in the production or introduction of, E of formula I. For example, if F₁ is a carboxylic acid, then -A-F₂ can represent the esterified form of the carboxylic acid. If F₁ is a cyano group, then -A-F₂ can represent, for example, a triazole ring with appended functionality for subsequent transformation. Although not depicted, the functionality -A-E may exist in commercially available starting materials, or be introduced in a single step to (i).

[0185] F_3 represents a functionality that is ultimately converted to, or used in the production or introduction of, W of formula I. For example, intermediate (ii) is brominated selectively to afford (iii) where F_3 is -Br. Using aryl-aryl coupling reactions known to one skilled in the art, W is introduced, preferably with functionality F_4 , into (iii) to make intermediate (v). F_4 represents a functionality that is ultimately converted to, or used in the production or introduction of, X of formula I. Alternatively, (iii) is converted to (iv) via introduction of E (for example an ester represented by -A- F_2 is converted to an amide represented by -A-E). Scheme 1 depicts only a subset of possible variations of making compounds of the invention.

Scheme 1



[0186] Still referring to Scheme 1, intermediate (v) can be converted to intermediate (vi) via introduction of E (intermediate (vi) can also be made from (iv) via introduction of -W- F_4). Also from intermediate (v), X is introduced, preferably with functionality F_5 , to make intermediate (vii). F_5 represents a functionality that is ultimately converted to, or used in the production or introduction of, Y of formula I. In many examples, -X- F_5 represents a linking group X with a reactive functionality, F_5 , that is used to attach Y to X. In such cases, (vii) is converted to (ix). Alternatively, intermediate (vii) is converted,

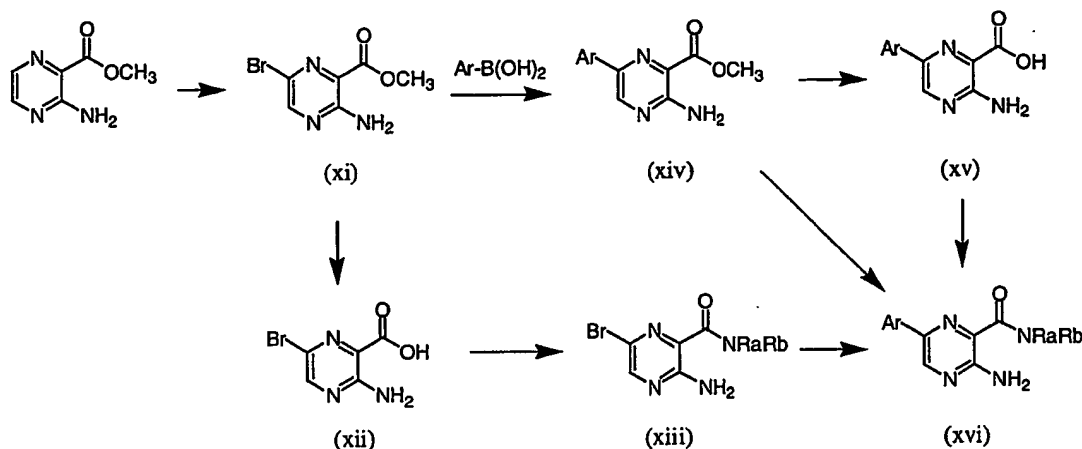
via introduction of $-X-F_5$, into (viii). Compounds of formula (x) are made from either intermediate (ix) or (viii). Compounds (x) are consistent with, and a subset of, formula I of the invention.

[0187] Thus compounds of the invention are generally made as depicted in Scheme 1. One skilled in the art would recognize that many variations of synthetic routes not included in Scheme 1 are possible, and thus are within the scope of the invention.

[0188] Scheme 2 depicts a general method of making 3-amino-pyrazine-2-carboxamides of the invention. Commercially available methyl 3-amino-pyrazine-2-carboxylate was brominated using the procedure of Russ, T., Ried, W., Ullrich, F., and Mutschler, E. *Arch. Pharm. (Weinheim)* 1992, 325, 761–767 (by analogy, bromination of commercially available 2-aminonicotinic acid (for example as described in United States Patents 3,950,160, and 4,361,700) affords 2-amino-5-bromopyridine-3-carboxylic acid. This intermediate can be used to make corresponding amino-pyridine analogues using similar methods similar to those described in Schemes 1 – 4 for 3-amino-pyrazine compounds.) The resulting bromo ester, (xi), was utilized, for example, in a Suzuki coupling with arylboronic acids by the method of W. J. Thompson, J. H. Jones, P. A. Lyle, and J. E. Thies *J. Org. Chem.* 1988, 53, 2052 – 2055 or a similar procedure. The resulting aryl substituted pyrazine ester, (xiv), was then hydrolyzed to the corresponding acid (xv). Acid (xv) was coupled with primary and secondary amines to give amides, (xvi), followed by removal of any protecting groups if necessary. Alternatively, aryl substituted pyrazine ester (xiv) was reacted directly with ammonia, methylamine, or other amines in a suitable solvent at elevated temperatures to afford amides (xvi). Again alternatively, bromopyrazine ester (xi) was converted (typically via hydrolysis) to bromopyrazine acid (xii), which was then coupled with amines to afford amides (xiii). The resulting bromopyrazine amides (xiii) were then, for example, coupled with arylboronic acids under Suzuki conditions to afford amides (xvi). In amides (xvi), the group $-NR_aR_b$ represents -E of formula I.

[0189] Alternatively, the ester moiety for example in intermediates (xi) and (xiv) were converted to an acid, amide, nitrile, aldehyde, hydroxymethyl, or other functional groups. These functional groups were then be elaborated into oximes, imines, hydrazones, or heterocycles such as triazoles, pyrimidines, oxadiazoles, and other heterocycles in accordance with group -A-E of formula I. More detailed descriptions of exemplary transformations are described below.

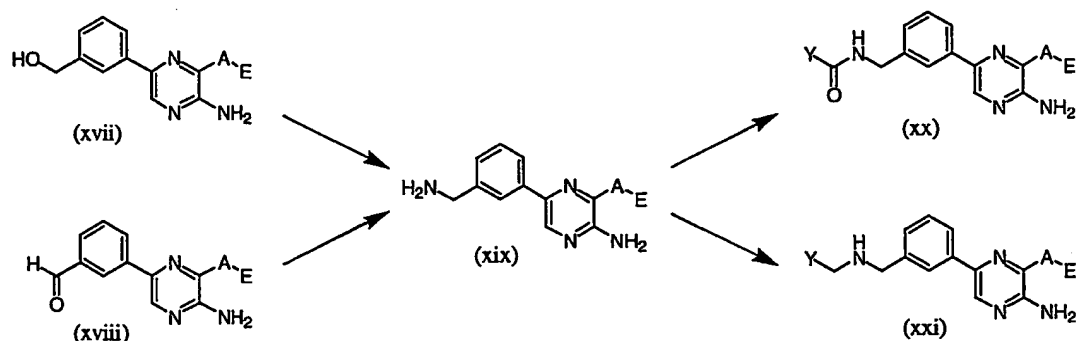
Scheme 2



[0190] Typically, but not necessarily, the aryl groups (denoted Ar in Scheme 2, corresponding to W in formula I) contained functional groups such as an ester, aldehyde, hydroxymethyl, amine, hydroxyl, or other groups that were modified or functionalized either before or after the the aryl groups were coupled to, for example the bromo pyrazine. Schemes 3 and 4 depict a number of typical transformations that were carried out to form group -X-Y onto W, in cases where W was a meta-phenylene. These are presented as examples and are not intended to limit the scope of the invention.

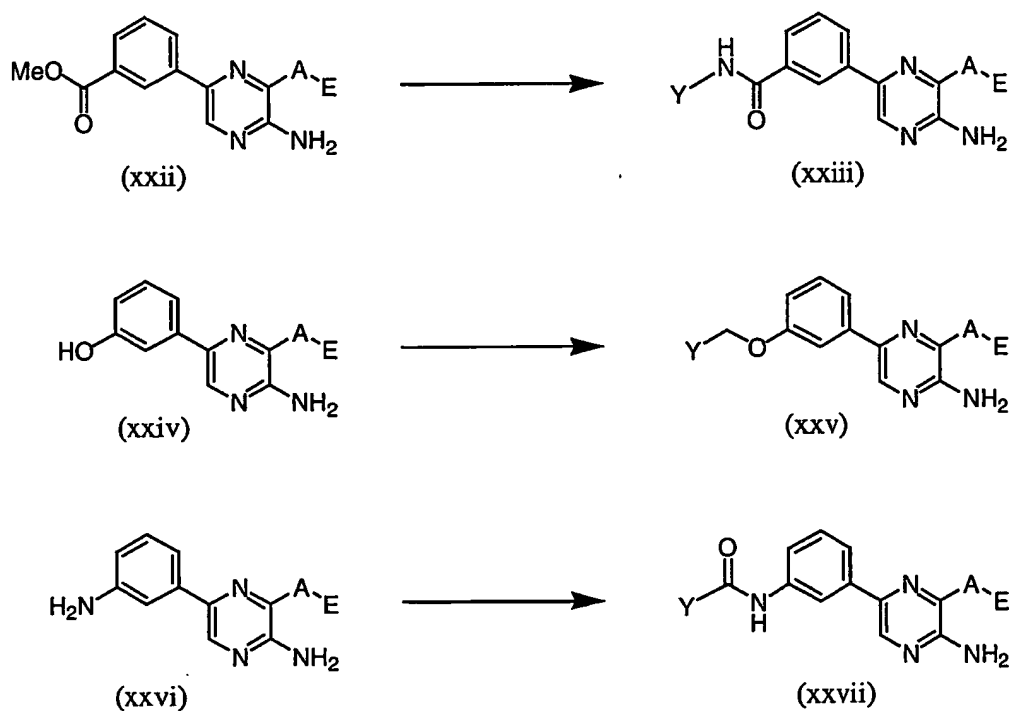
[0191] Referring to Scheme 3, benzylic amine intermediate (xix) was formed, for example, either from benzylic alcohol intermediate (xvii) or aryl aldehyde intermediate (xviii) using standard synthetic techniques. For example, the -OH group of (xvii) was transformed into a leaving group and subsequently displaced with ammonia or an ammonia equivalent (such as a phthalimide) to form (xix). Alternatively, aldehyde (xviii) is subjected to reductive amination conditions to form (xix). Intermediate (xix) was acylated, for example with acid chlorides, to form (xx) (according to formula I, W is meta-phenylene and X is -C(=O)NHCH₂-). Intermediate (xix) in many cases was, or otherwise could be, also reacted with sulfonyl chlorides, isocyanates, and other electrophiles. In another example, intermediate (xix) was subjected to reductive amination conditions to provide compounds (xxi).

Scheme 3



[0192] Scheme 4 shows additional transformations that were carried out to form group -X-Y onto W, in cases where W was a meta-phenylene. For example: esters (xxii) were converted to amides (xxiii), phenols (xxiv) were converted to ethers (xxv), and anilines (xxvi) were converted to amides (xxvii).

Scheme 4



Examples

[0193] The following examples serve to more fully describe the manner of using the above-described invention, as well as to set forth the best modes contemplated for

carrying out various aspects of the invention. It is understood that these examples in no way serve to limit the true scope of this invention, but rather are presented for illustrative purposes. All references cited herein are incorporated by reference in their entirety. Generally, but not necessarily, each example set out below describes a multi-step synthesis as outlined above.

Example 1

3-amino-6-phenyl-N-(phenylmethyl)pyrazine-2-carboxamide

[0194] Methyl 3-amino-6-bromo-pyrazine-2-carboxylate: Commercially available methyl 3-amino-pyrazine-2-carboxylate (available from Aldrich Chemical Company) was brominated using a procedure similar to that of Russ, T., Ried, W., Ullrich, F., and Mutschler, E. *Arch. Pharm. (Weinheim)* 1992, 325, 761 – 767. To a solution of methyl 3-amino-2-pyrazine carboxylate (30.0 g, 200 mmol) in acetic acid (200 mL), bromine (11 mL) was added slowly via addition funnel. After complete addition of bromine, sodium carbonate powder was added slowly until precipitation occurred. The reaction mixture was allowed to stir at room temperature for 2 h. The reaction mixture was reduced to half-volume in vacuo and then diluted with water (500 mL). The reaction mixture was shaken vigorously and the resulting solid was collected using vacuum filtration. The solid was washed with ether to afford a pure yellow solid (91 % yield).

[0195] Methyl 3-amino-6-phenylpyrazine-2-carboxylate: Methyl 3-amino-6-phenylpyrazine-2-carboxylate was prepared from methyl 3-amino-6-bromo-pyrazine-2-carboxylate using the procedure similar to that of W. J. Thompson, J. H. Jones, P. A. Lyle, and J. E. Thies *J. Org. Chem.* 1988, 53, 2052 – 2055.

[0196] 3-amino-6-phenylpyrazine-2- carboxylic acid: To a solution of methyl 3-amino-6-phenylpyrazine-2-carboxylate (1.2 g, 5.26 mmol) in methanol (5 mL) was added 6.0 N aqueous sodium hydroxide (6 mL) at room temperature. The solution was stirred at 50 C for 3 h. The reaction was neutralized slowly by adding 2.0 N hydrochloric acid until the pH of the solution became in the 2-3 range at 0° C. The solution was extracted with ethyl acetate (50 mL). The organic layer was washed with saturated aqueous sodium chloride (50 mL) and dried over magnesium sulfate. Filtration and concentration at reduced pressure afforded 3-amino-6-phenylpyrazine-2- carboxylic acid (0.59 g, 52% yield): MS (EI) for C₁₁H₉O₂: 216 (MH⁺)

[0197] 3-amino-6-phenyl-N-(phenylmethyl)pyrazine-2-carboxamide: To a solution of 3-amino-6-phenylpyrazine carboxylic acid (0.10 g, 0.47 mmol) in dichloromethane (3 mL) were added 1-ethyl-(3-dimethylaminopropyl)carbodiimide hydrochloride (EDCI, 0.11 g, 0.56 mmol), 1-hydroxybenzotriazole (HOBT, 75 mg, 0.56 mmol), and N-methylmorpholine (NMM, 56 mg, 0.56 mmol) at room temperature. The reaction was stirred for 15 min before benzylamine (0.56 mmol) was added. The reaction mixture was allowed to stir overnight. The reaction was diluted with ethyl acetate (200 mL) and washed with water (50 mL), saturated aqueous sodium bicarbonate (40 mL), 2.0 N aqueous hydrochloric acid (30 mL), and saturated aqueous sodium chloride (25 mL). The organic layer was dried over magnesium sulfate, filtered and concentrated at reduced pressure to afford product (91 mg, 67% yield) as light yellow solid: ^1H NMR (400 MHz, CDCl_3): δ 8.62 (s, 1H), 8.38 (br s, 1H), 7.84 (dd, 2H), 7.45 (m, 2H), 7.38 (m, 6H), 4.70 (d, 2H); MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_4\text{O}$: 305 (MH^+).

[0198] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0199] 3-amino-N,6-diphenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 9.92 (br s, 1H), 8.70 (s, 1H), 7.90 (dd, 2H), 7.72 (dd, 2H), 7.52 (m, 2H), 7.46 (m, 3H), 7.20 (t, 1H); MS (EI) for $\text{C}_{17}\text{H}_{14}\text{N}_4\text{O}$: 290 (MH^+).

[0200] 5-phenyl-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine: ^1H NMR (400 MHz, CDCl_3): δ 8.58 (s, 1H), 7.82(d, 2H), 7.45 (t, 2H), 7.40 (m, 1H), 6.40 (br s, 2H), 4.02 (m, 1H), 4.70(m, 1H), 2.04 (m, 4H); MS (EI) for $\text{C}_{15}\text{H}_{16}\text{N}_4\text{O}$: 269 (MH^+).

[0201] 3-amino-N-(cyclopropylmethyl)-6-phenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.62 (s, 1H), 8.10 (br s, 1H), 7.85 (dd, 2H), 7.50 (t, 2H), 7.40 (m, 1H), 3.30 (m, 2H), 1.10 (m, 1H), 0.80 (m, 2H), 0.30 (m, 2H); MS (EI) for $\text{C}_{15}\text{H}_{16}\text{N}_4\text{O}$: 269 (MH^+).

[0202] 3-amino-6-phenyl-N-(tetrahydrofuran-2-ylmethyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.60 (s, 1H), 8.30 (br s, 1H), 7.84 (d, 2H), 7.50 (t, 2H), 7.40 (m, 1H), 4.10 (m, 1H), 3.94 (m, 1H), 3.80 (m, 1H), 3.70 (m, 1H), 3.45 (m, 1H), 2.01 (m, 3H), 1.62 (m, 1H); MS (EI) for $\text{C}_{16}\text{H}_{18}\text{N}_4\text{O}_2$: 299 (MH^+).

[0203] 3-amino-N-cyclohexyl-6-phenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.80 (s, 1H), 7.92 (br s, 1H), 7.82 (d, 2H), 7.45 (t, 2H), 7.40 (m, 2H), 3.90 (m,

- 1H), 2.01 (m, 2H), 1.90 (m, 2H), 1.85(m, 1H), 1.40 (m, 5H); MS (EI) for C₁₇H₂₀N₄O: 297 (MH⁺).
- [0204] 3-amino-6-phenyl-N-(2-phenylethyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.82 (m, 2H), 8.06 (m, 2H), 7.65 (br s, 2H), 7.49 – 7.20 (m, 8H), 3.58 (t, 2H), 2.90 (t, 2H); MS (EI) for C₁₉H₁₈N₄O: 319.2 (MH⁺).
- [0205] 3-amino-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide: MS (EI) for C₂₀H₁₈N₄O: 331.2 (MH⁺).
- [0206] 3-amino-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.04 (d, 1H), 8.87 (s, 1H), 8.18 (m, 2H), 7.72 (br s, 2H), 7.49 – 7.18 (m, 7H), 5.60 (m, 1H), 3.02 (m, 1H), 2.86 (m, 1H), 2.44 (m, 1H), 2.21 (m, 1H); MS (EI) for C₂₀H₁₈N₄O: 331.2 (MH⁺).
- [0207] 3-amino-N-(2,3-dihydro-1H-inden-2-yl)-6-phenylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.04 (d, 1H), 8.87 (s, 1H), 8.18 (m, 2H), 7.72 (br s, 2H), 7.49 – 7.18 (m, 7H), 5.60 (m, 1H), 3.02 (m, 1H), 2.86 (m, 1H), 2.44 (m, 1H), 2.21 (m, 1H); MS (EI) for C₂₀H₁₈N₄O: 331.2 (MH⁺).
- [0208] 3-amino-N-cyclopentyl-6-phenylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.89 (d, 1H), 8.82 (s, 1H), 8.12 (m, 2H), 7.61 (br s, 2H), 7.46 – 7.15 (m, 7H), 4.79 (m, 1H), 3.28 (m, 2H), 3.11 (m, 2H); MS (EI) for C₁₆H₁₈N₄O: 283.2 (MH⁺).
- [0209] 3-amino-N-cyclopropyl-6-phenylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.83 (s, 1H), 8.65 (s, 1H), 8.13 (m, 2H), 7.65 (br s, 2H), 7.49 – 7.30 (m, 3H), 2.85 (m, 1H), 0.72 (m, 4H); MS (EI) for C₁₄H₁₄N₄O: 255.4 (MH⁺).

Example 2

3-Amino-6-naphthalen-1-ylpyrazine-2-carboxamide

- [0210] Methyl 3-amino-6-naphthalen-1-ylpyrazine-2-carboxylate: To a solution of methyl 3-amino-6-bromopyrazine-2-carboxylate (0.18 g, 0.74 mmol) in N,N-dimethylformamide (10 mL) were added [1,1'-bis (diphenylphosphino)ferrocene]dichloropalladium(II) complex with dichloromethane (1:1, 60 mg, 0.07 mmol), 1-naphthylboronic acid (0.32 g, 1.89 mmol), and triethylamine (0.15g, 1.52 mmol). The solution was degassed with nitrogen for 3-5 min. The reaction was then heated to 85-90° C overnight. The reaction was cooled to room temperature and diluted with ethyl acetate (300 mL). The crude

solution was passed through a pad of silica gel under vacuum. The filtrate was washed with saturated aqueous sodium bicarbonate (50 mL) and saturated aqueous sodium chloride (50 mL). The organic layer was dried over magnesium sulfate, filtered and concentrated at reduced pressure to afford impure product. Column purification on silica (8:2 hexanes/ ethyl acetate) afforded methyl 3-amino-6-naphthalen-1-ylpyrazine-2-carboxylate (80 mg, 39% yield) as yellow solid: ^1H NMR (400 MHz, CDCl_3): δ 8.52 (br s, 1H), 8.12 (br s, 1H), 7.92 (m, 2H), 7.52 (m, 5H), 4.02 (s, 3H); MS (EI) for $\text{C}_{16}\text{H}_{13}\text{N}_3\text{O}_2$: 280 (MH^+).

[0211] 3-amino-6-naphthalen-1-ylpyrazine-2-carboxamide: A solution of methyl 3-amino-6-naphthalen-1-ylpyrazine-2-carboxylate (80 mg, 0.29 mmol) in methanol (30 mL) was saturated with ammonia gas at 0°C in a high-pressure test tube. The reaction tube was tightly sealed and heated to $80\text{--}90^\circ\text{C}$ for 4h. Reaction cooled to room temperature and evaporated at reduced pressure to afford 3-amino-6-naphthalen-1-ylpyrazine-2-carboxamide (59 mg, 79% yield) as yellow product: ^1H NMR (400 MHz, CDCl_3): δ 8.80 (s, 1H), 8.10 (d, 1H), 7.90 (d, 2H), 7.58 (m, 6H); MS (EI) for $\text{C}_{15}\text{H}_{12}\text{N}_4\text{O}$: 265 (MH^+).

[0212] 3-amino-6-(3-chlorophenyl)pyrazine-2-carboxamide: To a 15 mL sealed tube was added methyl 3-amino-6-(3-chlorophenyl)pyrazine-2-carboxylate (65 mg, 0.246 mmol) and methanol (8 mL). The suspension was cooled to 0°C and NH_3 (g) was bubbled until saturation was achieved. The tube was sealed and heated at 85°C overnight with stirring. During initial heating, the reaction mixture became homogeneous. The tube was removed from the heat and allowed to cool to room temperature. Upon cooling a yellow, needle-like precipitate formed. The precipitate was collected on filter paper to give 3-amino-6-(3-chlorophenyl)pyrazine-2-carboxamide (53.9 mg, 88% yield) as a yellow solid: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 8.88 (s, 1H), 8.42 (br s, 2H), 8.30 (t, 1H), 8.11 (d, 1H), 7.69 (br s, 2H), 7.46-7.40 (m, 2H); MS (EI) for $\text{C}_{11}\text{H}_9\text{N}_4\text{OCl}$: 249 (MH^+).

[0213] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0214] 3-amino-6-naphthalen-2-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 9.02 (s, 1H), 8.80 (s, 1H), 8.40 (m, 2H), 8.02 (m, 3H), 7.80 (br s, 1H), 7.50 (t, 2H); MS (EI) for $\text{C}_{15}\text{H}_{12}\text{N}_4\text{O}$: 265 (MH^+).

- [0215] 3-amino-6-biphenyl-4-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.90 (s, 1H), 8.38 (br s, 1H), 8.30 (d, 2H), 7.70 (m, 5H), 7.50 (m, 2H), 7.34 (m, 1H); MS (EI) for $\text{C}_{17}\text{H}_{14}\text{N}_4\text{O}$: 291 (MH^+).
- [0216] 3-amino-6-(1-benzofuran-2-yl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.85 (s, 1H), 8.60 (br s, 1H), 7.80 (s, 1H), 7.60 (m, 3H), 7.30 (m, 2H); MS (EI) for $\text{C}_{13}\text{H}_{10}\text{N}_4\text{O}$: 255 (MH^+).
- [0217] 3-amino-6-biphenyl-3-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.85 (d, 2H), 7.65 (t, 3H), 7.55 (t, 1H), 7.45 (t, 3H), 7.40 (t, 1H); MS (EI) for $\text{C}_{17}\text{H}_{14}\text{N}_4\text{O}$: 291 (MH^+).
- [0218] 3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoic acid: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.95 (s, 1H), 8.80 (br s, 2H), 8.50 (s, 1H), 8.30 (d, 1H), 7.98 (d, 1H), 7.60 (br t, 1H); MS (EI) for $\text{C}_{12}\text{H}_{10}\text{N}_4\text{O}_3$: 259 (MH^+).
- [0219] 3-amino-6-{4-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (s, 1H), 8.30 (br s, 1H), 8.10 (d, 2H), 7.65 (br s, 1H), 7.49 (d, 2H), 7.40 (t, 2H), 7.35 (t, 1H), 7.10 (d, 2H), 5.20 (s, 2H); MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_4\text{O}_2$: 321 (MH^+).
- [0220] 3-amino-6-(2-chlorophenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.53 (s, 1H), 7.76-7.73 (m, 1H), 7.59-7.56 (m, 1H), 7.46-7.43 (m, 2H); MS (EI) for $\text{C}_{11}\text{H}_9\text{N}_4\text{OCl}$: 249 (MH^+).
- [0221] 3-amino-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.82 (s, 1H), 7.69-7.65 (m, 4H), 7.33 (t, 1H), 6.91 (dd, 1H), 3.82 (s, 3H); MS (EI) for $\text{C}_{12}\text{H}_{12}\text{N}_4\text{O}_2$: 245 (MH^+).
- [0222] 3-amino-6-(2-methylphenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.38 (s, 1H), 7.48-7.47 (m, 1H), 7.30-7.28 (m, 3H); MS (EI) for $\text{C}_{12}\text{H}_{12}\text{N}_4\text{O}$: 229 (MH^+).
- [0223] 3-amino-6-[2-(methyloxy)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.72 (s, 1H), 7.94-7.91 (m, 1H), 7.36-7.34 (m, 1H), 7.14-7.12 (m, 1H), 7.07-7.05 (m, 1H), 3.86 (s, 3H); MS (EI) for $\text{C}_{12}\text{H}_{12}\text{N}_4\text{O}_2$: 245 (MH^+).
- [0224] 3-amino-6-(2,4-difluorophenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.59 (s, 1H), 8.28-8.26 (m, 2H), 7.17 (br s, 2H), 7.38-7.37 (m, 1H), 7.20 (m, 1H); MS (EI) for $\text{C}_{11}\text{H}_8\text{N}_4\text{OF}_2$: 251 (MH^+).

- [0225] 3-amino-6-(3-fluorophenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d6-DMSO): δ 8.89 (s, 1H), 8.11 (d, 1H), 7.98 (d, 1H), 7.48-7.46 (m, 1H), 7.17-7.16 (m, 1H); MS (EI) for $\text{C}_{11}\text{H}_9\text{N}_4\text{OF}$: 233 (MH^+).
- [0226] 3-amino-6-(3,4-difluorophenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d6-DMSO): δ 8.88 (s, 1H), 8.41-8.35 (m, 1H), 8.02 (m, 1H), 7.52-7.45 (m, 1H); MS (EI) for $\text{C}_{11}\text{H}_8\text{N}_4\text{OF}_2$: 251 (MH^+).
- [0227] 3-amino-6-(4-fluorophenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d6-DMSO): δ 8.84 (m, 2H), 8.25-8.22 (m, 2H), 7.31-7.26 (m, 2H), 2.84 (d, 3H); MS (EI) for $\text{C}_{12}\text{H}_{11}\text{N}_4\text{OF}$: 247 (MH^+).
- [0228] 3-amino-6-{3-[(trifluoromethyl)oxy]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d6-DMSO): δ 8.91 (s, 1H), 8.23 (s, 1H), 8.20-8.18 (m, 1H), 7.59-7.54 (t, 1H), 7.35-7.33 (d, 1H); MS (EI) for $\text{C}_{12}\text{H}_9\text{N}_4\text{O}_2\text{F}_3$: 299 (MH^+).
- [0229] 3-amino-6-(3-methylphenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d6-DMSO): δ 8.82 (s, 1H), 8.00 (s, 1H), 7.93-7.91 (d, 1H), 7.34-7.31 (t, 1H), 7.18-7.16 (m, 1H), 2.38 (s, 3H); MS (EI) for $\text{C}_{12}\text{H}_{12}\text{N}_4\text{O}$: 229 (MH^+).
- [0230] 3-amino-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d6-DMSO): δ 8.84 (s, 1H), 7.78-7.77 (m, 1H), 7.71-7.70 (m, 2H), 7.50-7.59 (m, 2H), 7.43-7.39 (m, 2H), 7.38-7.31 (m, 2H), 7.02-6.99 (m, 1H), 5.21 (s, 2H); MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_4\text{O}_2$: 321 (MH^+).
- [0231] 3-amino-6-(3-hydroxyphenyl)pyrazine-2-carboxamide: To a 25 mL recovery flask was added 3-amino-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide (33 mg, 0.103 mmol) and ethyl acetate (10 mL). A catalytic amount of 10% Pd/C was added. The flask was sealed with a septum and fitted with an H_2 balloon. The reaction mixture was stirred overnight at room temperature. The reaction mixture was passed through a Celite pad. The filtrate was concentrated to give 3-amino-6-(3-hydroxyphenyl)pyrazine-2-carboxamide (23 mg, 96% yield) as a yellow powder: ^1H NMR (400 MHz, d6-DMSO): δ 9.47 (s, 1H), 8.74 (s, 1H), 7.55-7.53 (m, 1H), 7.46 (m, 1H), 7.25-7.22 (t, 1H), 6.78-6.76 (m, 1H); MS (EI) for $\text{C}_{11}\text{H}_{10}\text{N}_4\text{O}_2$: 231 (MH^+).
- [0232] 3-amino-N,N-dimethyl-6-phenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.54 (s, 1H), 7.89 (m, 2H), 7.48 – 7.33 (m, 3H), 5.88 (br s, 2H), 3.33 (s, 3H), 3.17 (s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_4\text{O}$: 243.1 (MH^+).

- [0233] 3-amino-6-[4-(ethyloxy)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.78 (s, 1H), 8.28 (s, 1H), 8.08 (m, 2H), 7.66 (s, 1H), 7.56 (br s, 2H), 6.98 (m, 2H), 4.08 (q, 2H), 1.35 (t, 3H); MS (EI) for $\text{C}_{31}\text{H}_{14}\text{N}_4\text{O}_2$: 259.4 (MH^+).
- [0234] 3-amino-6-[3-(ethyloxy)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.85 (s, 1H), 8.30 (s, 1H), 7.96 – 7.28 (m, 6H), 6.92 (m, 1H), 4.13 (q, 2H), 1.35 (t, 3H); MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_4\text{O}_2$: 259.4 (MH^+).
- [0235] 3-amino-6-(1,3-benzodioxol-5-yl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.78 (s, 1H), 8.36 (s, 1H), 7.85 (s, 1H), 7.79 – 7.37 (m, 4H), 6.96 (d, 1H), 6.07 (s, 2H); MS (EI) for $\text{C}_{12}\text{H}_{10}\text{N}_4\text{O}_3$: 259.4 (MH^+).
- [0236] 3-amino-6-[3-(aminocarbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.92 (s, 1H), 8.54 (s, 1H), 8.40 (s, 1H), 8.32 (d, 1H), 8.16 (s, 1H), 7.95 – 7.42 (m, 6H); MS (EI) for $\text{C}_{12}\text{H}_{11}\text{N}_5\text{O}_2$: 258.4 (MH^+).
- [0237] 6-[3-(acetilamino)phenyl]-3-aminopyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.99 (s, 1H), 8.72 (s, 1H), 8.11 (m, 2H), 7.94 – 7.46 (m, 5H), 7.37 (m, 1H), 2.06 (s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{13}\text{N}_5\text{O}_2$: 272.4 (MH^+).
- [0238] 3-amino-6-[4-(dimethylamino)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.73 (s, 1H), 8.21 (s, 1H), 7.97 (m, 2H), 7.64 (s, 1H), 7.45 (br s, 2H), 6.76 (m, 2H), 2.95 (s, 6H); MS (EI) for $\text{C}_{13}\text{H}_{15}\text{N}_5\text{O}$: 258.4 (MH^+).
- [0239] methyl 3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoate: ^1H NMR (400 MHz, d_6 -DMSO): 8.90 (s, 1H), 8.57 (s, 1H), 8.48 (m, 1H), 8.31 (s, 1H), 8.00 – 7.42 (m, 5H), 3.90 (s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{12}\text{N}_4\text{O}_3$: 273.4 (MH^+).
- [0240] 3-amino-N-methyl-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.82 (m, 2H), 7.92 – 7.19 (m, 10H), 7.03 (m, 1H), 5.22 (s, 2H), 2.85 (s, 3H); MS (EI) for $\text{C}_{19}\text{H}_{18}\text{N}_4\text{O}_2$: 335.4 (MH^+).
- [0241] 3-amino-6-(3-hydroxyphenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.49 (s, 1H), 8.73 (m, 2H), 7.87 – 7.34 (m, 4H), 7.25 (m, 1H), 6.79 (m, 1H), 2.85 (d, 3H); MS (EI) for $\text{C}_{12}\text{H}_{12}\text{N}_4\text{O}_2$: 245.4 (MH^+).
- [0242] 3-amino-N-[2-(methyloxy)ethyl]-6-phenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.85 (m, 2H), 8.15 (m, 2H), 7.68 (br s, 2H), 7.47 (m, 2H), 7.40 (m, 1H), 3.50 (m, 4H), 3.29 (s, 3H); MS (EI) for $\text{C}_{14}\text{H}_{16}\text{N}_4\text{O}_2$: 273.4 (MH^+).

- [0243] N-[2-(acetylamino)ethyl]-3-amino-6-phenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 8.94 (t, 1H), 8.86 (s, 1H), 8.19 (m, 2H), 8.05 (t, 1H), 7.66 (br s, 1H), 7.46 (m, 2H), 7.38 (m, 1H), 3.38 (m, 2H), 3.27 (m, 2H), 1.83 (s, 3H); MS (EI) for $\text{C}_{15}\text{H}_{17}\text{N}_5\text{O}_2$: 300.4 (MH^+).
- [0244] 3-amino-6-phenylpyrazine-2-carbohydrazide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 10.02 (s, 1H), 8.82 (s, 1H), 8.19 (m, 2H), 7.58 (br s, 2H), 7.43 (m, 2H), 7.35 (m, 2H), 7.32 (m, 1H), 4.57 (m, 2H); MS (EI) for $\text{C}_{11}\text{H}_{11}\text{N}_5\text{O}$: 230.3 (MH^+).
- [0245] 3-amino-N-hydroxy-6-phenylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 11.42 (s, 1H), 9.12 (s, 1H), 8.83 (s, 1H), 8.20 (m, 2H), 7.53 (br s, 2H), 7.42 (m, 2H), 7.36 (m, 1H); MS (EI) for $\text{C}_{11}\text{H}_{10}\text{N}_4\text{O}_2$: 231.3 (MH^+).
- [0246] 3-amino-N-(2-hydroxyethyl)-6-(3-methylphenyl)pyrazine-2-carboxamide: A solution of 3-amino-6-bromo-2-carbomethoxypyrazine (100 mg, 0.4 mmol) and 2-aminoethanol (500 mg, 8.2 mmol) in THF (5 mL) was heated at reflux for 2 days. The reaction mixture was cooled, diluted with ethyl acetate (20 mL), washed with water then brine, dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give 3-amino-N-(2-hydroxyethyl)-6-bromo-pyrazine-2-carboxamide, a pale yellow oil, (108 mg, 96% yield) MS (EI) for $\text{C}_7\text{H}_9\text{BrN}_4\text{O}_2$: 262 (MH^+).
- [0247] The intermediate 3-amino-N-(2-hydroxyethyl)-6-bromopyrazine-2-carboxamide (108 mg, 0.4 mmole), [1,1'-bis-(diphenylphosphino)ferrocene]dichloropalladium(II)-dichloromethane complex (19 mg, 0.04 mmol), 3-methylphenylboronic acid (67 mg, 0.5 mmol) in DMF (2 mL) was de-gassed with nitrogen. Triethylamine (0.12 mL, 0.8 mmol) was then added and the resulting mixture was heated at 85 °C for 2.5 hours. The reaction mixture was cooled to room temperature, diluted with ethyl acetate (10 mL), filtered over silica gel, washed with saturated sodium bicarbonate solution (10 mL), dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give methyl 3-amino-N-(2-hydroxyethyl)-6-(3-methylphenyl)pyrazine-2-carboxamide as a solid: (95 mg, 84% yield): ^1H NMR (400 MHz, DMSO-d_6): 8.78 (s, 1H), 8.78-8.61 (b, 1H), 7.89 (s, 1H), 7.49-7.52 (t, 1H), 7.51-7.49 (d, 1H), 7.14-7.12 (d, 1H), 3.50-3.35 (m, 2H), 2.89-7.0 (m, 2H), 2.51 (s, 3H); MS (EI) for $\text{C}_{14}\text{H}_{16}\text{N}_4\text{O}_2$: 273 (MH^+).

Example 3**3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide**

[0248] 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide: An ice-cooled solution of 50 g of methyl 3-amino-6-bromo-pyrazine-2-carboxylate in 500 mL of MeOH was saturated with methylamine gas. The reaction was then heated at 85 °C in a sealed, stainless steel Parr pressure vessel for 2 h, and cooled to room temperature, and then on an ice water bath. The mixture was concentrated to an oil on a rotary evaporator under reduced pressure. The crude oil was dissolved in EtOAc and washed with water, dried (Na₂SO₄), filtered, and concentrated to give 43.9 g (88.0% yield) of product as a solid.

[0249] 3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide: A suspension of 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide (69.3mg, 0.3 mmol), [4-(tetrahydro-2H-pyran-2-yloxy)phenyl boronic acid (111 mg, 0.5 mmol) [1,1-bis(diphenylphosphino)-ferrocene]dichloropalladium(II) complex with dichloromethane (1:1) (20.4 mg, 0.025 mmol), and triethylamine (121 mg, 1.2 mmol) in DMF (20 mL) was stirred at 80 °C under nitrogen for 12 h. DMF was removed under vacuo. The residue was extracted with ethyl acetate (3 x 50 mL). The organic layers were washed with brine (10 mL) then dried over anhydrous sodium sulfate. Filtration, concentration and column chromatography on silica gave 3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide (68.3 mg, 70% yield) as a yellow solid. ¹H NMR (400 MHz, d₆-DMSO): 8.8 (m, 2H), 8.1 (d, 2H), 7.1 (d, 2H), 5.56 (s, 1H), 3.78 (m, 2H), 3.45 (m, 2H), 1.98- 1.43 (m, 6H); MS (EI) for C₁₇H₂₀N₄O₃: 329 (MH⁺).

[0250] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0251] 3-amino-6-[4-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, CDCl₃): δ 8.52 (s, 1H), 8.02 (br s, 1H), 7.80 (d, 2H), 7.00 (br d, 2H), 7.55 (t, 2H), 4.12 (q, 2H), 3.02 (d, 3H), 1.43 (t, 3H); MS (EI) for C₁₄H₁₆N₄O₂: 273 (MH⁺).

[0252] 3-amino-N-methyl-6-(3-methylphenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, CDCl₃): δ 8.60 (s, 1H), 8.01 (br s, 1H), 7.64 (br d, 2H), 7.38 (t, 1H), 7.25 (br d, 1H), 3.02 (d, 3H), 2.42 (s, 3H); MS (EI) for C₁₃H₁₄N₄O: 243 (MH⁺).

- [0253] 3-amino-N-methyl-6-[4-(methyloxy)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.57 (s, 1H), 8.02 (br s, 1H), 7.80 (br d, 2H), 7.02 (br d, 2H), 3.84 (s, 3H), 3.10 (d, 3H); MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_4\text{O}_2$: 259 (MH^+).
- [0254] 3-amino-N-methyl-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.60 (s, 1H), 8.02 (br s, 1H), 7.40 (m, 3H), 6.90 (br d, 1H), 3.92 (s, 3H), 3.02 (br s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_4\text{O}_2$: 259 (MH^+).
- [0255] 3-amino-6-(4-fluoro-3-methylphenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.52 (s, 1H), 7.94 (br s, 1H), 7.65 (m, 2H), 7.12 (t, 1H), 3.02 (d, 3H), 2.40 (s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{13}\text{N}_4\text{O F}$: 261 (MH^+).
- [0256] 3-amino-N-methyl-6-pyridin-3-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.40 (br s, 1H), 8.94 (br s, 2H), 8.54 (m, 2H), 7.44 (t, 1H), 2.90 (br s, 3H); MS (EI) for $\text{C}_{11}\text{H}_{11}\text{N}_5\text{O}$: 230 (MH^+).
- [0257] 3-amino-6-[4-(dimethylamino)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.58 (s, 1H), 8.12 (br s, 1H), 7.75 (br d, 2H), 6.80 (br d, 2H), 3.02 (m, 9H); MS (EI) for $\text{C}_{14}\text{H}_{17}\text{N}_5\text{O}$: 272 (MH^+).
- [0258] 3-amino-N-methyl-6-[4-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.80 (br s, 1H), 8.80 (br s, 2H), 8.15 (br d, 2H), 7.30 (br d, 2H), 3.02 (br s, 3H), 2.80 (br s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{15}\text{N}_5\text{O}_3\text{S}$: 322 (MH^+).
- [0259] 3-amino-6-(3,5-dimethylphenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.60 (s, 1H), 8.02 (br s, 1H), 7.45 (br d, 2H), 7.05 (br s, 1H), 3.02 (br s, 3H), 2.40 (br s, 6H); MS (EI) for $\text{C}_{14}\text{H}_{16}\text{N}_4\text{O}$: 257 (MH^+).
- [0260] 6-(4-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.70 (br s, 1H), 8.05 (br d, 2H), 7.95 (br d, 3H), 3.10 (d, 3H), 2.60 (s, 3H); MS (EI) for $\text{C}_{14}\text{H}_{14}\text{N}_4\text{O}_2$: 271 (MH^+).
- [0261] 3-amino-6-[3,4-bis(methyloxy)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.58 (s, 1H), 7.95 (br s, 1H), 7.40 (dd, 1H), 7.36 (d, 1H), 6.82 (d, 1H), 4.02 (s, 3H), 3.92 (s, 3H), 3.02 (d, 3H); MS (EI) for $\text{C}_{14}\text{H}_{16}\text{N}_4\text{O}_3$: 289 (MH^+).
- [0262] 3-amino-6-(3-cyanophenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.89 (m, 2H), 8.70 (br s, 1H), 8.45 (d, 1H), 7.80 (d, 1H), 7.60 (t, 1H), 2.82 (br s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{11}\text{N}_5\text{O}$: 254 (MH^+).

- [0263] 3-amino-N-methyl-6-[3-(1H-tetrazol-5-yl)phenyl]pyrazine-2-carboxamide: To a solution of 3-amino-6-(3-cyanophenyl)-N-methylpyrazine-2-carboxamide (93 mg, 0.37 mmol) in dry N,N-dimethylformamide (1 mL) were added sodium azide (27 mg, 0.41 mmol), and ammonium chloride (22 mg, 0.41 mmol). The reaction was heated to 100-110° C overnight. Reaction cooled to room temperature and treated with saturated aqueous sodium bicarbonate (20 mL). The aqueous layer was washed with ethyl acetate (2X30 mL). The sodium bicarbonate layer was cooled to 0° C and neutralized with 2.0 N hydrochloric acid until the pH of the solution became in the 2-3 range. The aqueous layer was extracted with ethyl acetate (100 mL) and washed with water (20 mL) and saturated aqueous sodium chloride (30 mL), dried over magnesium sulfate, filtered, and concentrated at reduced pressure to afford 3-amino-N-methyl-6-[3-(1H-tetrazol-5-yl)phenyl]pyrazine-2-carboxamide (25 mg, 23% yield) as light yellow solid. ¹H NMR (400 MHz, d₆-DMSO): 8.92 (s, 1H), 8.84 (m, 1H), 8.70 (br s, 1H), 8.40 (br d, 1H), 8.12 (br d, 1H), 7.70 (t, 1H), 2.92 (br s, 3H); MS (EI) for C₁₃H₁₂N₈O: 297 (MH⁺).
- [0264] 3-amino-6-[3-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.83 (s, 1H), 8.81 (m, 1H), 7.70 (m, 2H), 7.62 (br s, 2H), 7.36 (m, 1H), 6.93 (m, 1H), 4.12 (q, 2H), 2.85 (d, 3H), 1.36 (t, 3H); MS (EI) for C₁₄H₁₆N₄O₂: 273.4 (MH⁺).
- [0265] 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.66 (m, 2H), 7.58 (br s, 2H), 7.27 (m, 2H), 7.11 (m, 1H), 6.59 (m, 1H), 5.09 (s, 2H), 2.85 (d, 3H); MS (EI) for C₁₂H₁₃N₅O: 244.4 (MH⁺).
- [0266] 6-[3-(acetylamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 10.01 (s, 1H), 8.69 (m, 2H), 8.03 (m, 1H), 7.83 (m, 1H), 7.78 (m, 1H), 7.63 (br s, 2H), 7.38 (m, 1H), 2.85 (d, 3H), 2.08 (s, 3H); MS (EI) for C₁₂H₁₃N₅O: 286.4 (MH⁺).
- [0267] 3-amino-6-[3-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.82 (s, 1H), 8.78 (m, 1H), 8.04 (m, 2H), 7.64 (br s, 2H), 7.42 (m, 1H), 7.35 (m, 1H), 5.23 (t, 1H), 4.58 (d, 2H), 2.86 (d, 3H); MS (EI) for C₁₃H₁₄N₄O₂: 259.4 (MH⁺).
- [0268] 3-amino-6-[4-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.84 (s, 1H), 8.82 (m, 1H), 8.14 (d, 2H), 7.62 (br s, 2H),

- 7.40 (d, 2H), 5.24 (t, 1H), 4.55 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{13}H_{14}N_4O_2$: 259.4 (MH^+).
- [0269] 3-amino-N-methyl-6-{3-[(1E)-N-methylethanimidoyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.86 (s, 1H), 8.81 (m, 1H), 8.35 (m, 1H), 8.22 (m, 1H), 7.80 (m, 1H), 7.68 (br s, 2H), 7.48 (t, 1H), 3.29 (s, 3H), 2.85 (d, 3H), 2.30 (s, 3H); MS (EI) for $C_{15}H_{17}N_5O$: 284.4 (MH^+).
- [0270] 6-[4-(acetylamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 10.06 (s, 1H), 8.81 (m, 1H), 8.80 (s, 1H), 8.11 (d, 2H), 7.67 (d, 2H), 7.60 (br s, 2H), 2.84 (d, 3H), 2.07 (s, 3H); MS (EI) for $C_{14}H_{15}N_5O_2$: 286.4 (MH^+).
- [0271] 3-amino-N-methyl-6-{3-[(methylamino)carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.90 (s, 1H), 8.84 (m, 1H), 8.54 (m, 1H), 8.47 ((s, 1H), 8.34 (d, 1H), 7.82 (d, 1H), 7.72 (br s, 2H), 7.56 (t, 1H), 2.84 (d, 3H), 2.82 (d, 3H); MS (EI) for $C_{14}H_{15}N_5O_2$: 286.4 (MH^+).
- [0272] 6-(3-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.94 (s, 1H), 8.87 (m, 1H), 8.57 (t, 1H), 8.48 (m, 1H), 7.93 (m, 1H), 7.75 (br s, 2H), 7.62 (t, 1H), 2.86 (d, 3H), 2.70 (s, 3H); MS (EI) for $C_{14}H_{14}N_4O_2$: 271.4 (MH^+).
- [0273] 3-amino-N-methyl-6-(3-[[phenylmethyl]amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.14 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.54 (t, 1H), 8.39 (m, 1H), 7.89 (m, 1H), 7.70 (br s, 2H), 7.58 (t, 1H), 7.38 – 7.32 (m, 4H), 7.24 (m, 1H), 4.54 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{19}N_5O_2$: 362.1 (MH^+).
- [0274] 3-amino-6-[3-(aminocarbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO/ CD_3OD): 8.90 (m, 1H), 8.57 (m, 1H), 8.32 (m, 1H), 7.91 (m, 1H), 7.57 (m, 1H), 2.90 (m, 3H); MS (EI) for $C_{13}H_{13}N_5O_2$: 272.0 (MH^+).
- [0275] 3-amino-6-{3-[(dimethylamino)carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.86 (m, 2H), 8.20 (m, 2H), 7.70 (br s, 2H), 7.49 (m, 1H), 7.34 (m, 1H), 3.02 (s, 3H), 2.92 (s, 3H), 2.83 (m, 3H); MS (EI) for $C_{15}H_{17}N_5O_2$: 300.4 (MH^+).
- [0276] 3-amino-N-methyl-6-[4-(methylsulfonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.95 (m, 2H), 8.45 (m, 2H), 8.11 (br s, 2H), 7.96 (m, 2H), 3.26 (s, 3H), 2.85 (m, 3H); MS (EI) for $C_{13}H_{14}N_4O_3S$: 307.4 (MH^+).

- [0277] 3-amino-6-(3-aminophenyl)-N-ethylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 8.71 (m, 1H), 8.65 (s, 1H), 7.60 (br s, 2H), 7.27 (m, 2H), 7.11 (m, 1H), 6.59 (m, 1H), 5.10 (s, 2H), 3.34 (m, 2H), 1.16 (m, 3H); MS (EI) for $\text{C}_{13}\text{H}_{15}\text{N}_5\text{O}$: 258.3 (MH^+).
- [0278] 3-amino-N-methyl-6-pyrimidin-5-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 9.57 (m, 2H), 9.15 (s, 1H), 8.98 (m, 2H), 8.94 (s, 1H), 2.8 (d, 3H), MS (EI) for $\text{C}_{10}\text{H}_{10}\text{N}_6\text{O}_2$: 231 (MH^+).
- [0279] 3-amino-N-methyl-6-[3-(methylsulfonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 8.98 (s, 1H), 8.84 (m, NH), 8.52 (m, 2H), 7.85 (d, 1H), 7.74 (t, 1H), 3.15 (s, 3H); 2.85 (d, 3H); MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_4\text{O}_3\text{S}$: 307 (MH^+).
- [0280] 3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide: Prepared as above.
- [0281] 3-amino-N-methyl-6-(4-hydroxyoxyphenyl)pyrazine-2-carboxamide: 3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide (63 mg) was suspended in methanol followed by addition of aqueous hydrochloric acid (2 mL). The mixture was stirred for 15 min., poured into water (10 mL), and extracted with ethyl acetate (10 mL). The organic layer was washed with saturate aqueous sodium chloride then dried over anhydrous sodium sulfate. Filtration, concentration and column chromatography on silica followed by drying in vacuo afforded 3-amino-N-methyl-6-(4-hydroxyoxyphenyl)pyrazine-2-carboxamide (30 mg, 64% yield): ^1H NMR (400 MHz; DMSO-D_6): 9.62 (s, 1H); 8.78 (m, 1H); 8.82 (m, 1H); 8.7 (s, 1H); 7.98 (d, 2H); 7.5 (br s, 1H); 6.8 (d, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{12}\text{H}_{12}\text{N}_4\text{O}_2$: 245 (MH^+).
- [0282] 3-amino-N-methyl-6-(3-thienyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO-D_6): 8.85 (m, 1H); 8.76 (s, 1H); 8.25 (m, 1H); 7.92 (d, 1H); 7.64 (m, 1H); 2.82 (d, 3H); MS (EI) for $\text{C}_{10}\text{H}_{10}\text{N}_4\text{OS}$: 235 (MH^+).
- [0283] 3-amino-6-(1H-indol-5-yl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO-D_6): 11.2 (s, 1H); 8.8 (s, 1H); 8.75 (m, 1H); 8.3 (s, 1H); 7.9 (d, 1H); 7.4 (d, 1H); 7.35 (t, 1H); 6.45 (s, 1H); 2.85 (d, 3H); MS (EI) for $\text{C}_{14}\text{H}_{13}\text{N}_5\text{O}$: 268 (MH^+).
- [0284] 3-amino-N-methyl-6-(2-thienyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO-D_6): 8.7 (s, 1H); 8.48 (m, 1H); 7.75 (d, 1H); 7.62 (br s, 2H); 7.57 (d, 1H); 7.15 (t, 1H); 2.85 (d, 3H); MS (EI) for $\text{C}_{10}\text{H}_{10}\text{N}_4\text{OS}$: 235 (MH^+).

- [0285] 3-amino-N-methyl-6-pyridin-4-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz; CD_3OD): 8.82 (s, 1H); 8.55 (d, 2H); 8.1 (d, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{11}\text{H}_{11}\text{N}_5\text{O}$: 230 (MH^+).
- [0286] 3-amino-N-methyl-6-naphthalen-2-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 8.99 (s, 1H), 8.89 (m, 1H), 8.69 (s, 1H), 8.38 (d, 1H), 7.98 (d, 2H), 7.92 (d, 1H), 7.74 (br s, 2H), 7.52 (m, 2H), 2.88 (d, 3H); MS (EI) for $\text{C}_{16}\text{H}_{14}\text{N}_4\text{O}$: 279.2 (MH^+).
- [0287] 3-amino-N-methyl-6-quinolin-3-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 9.76 (d, 1H), 9.05 (s, 2H), 8.99 (m, 1H), 8.02 (m, 2H), 7.89 (br s, 2H), 7.75 (m, 1H), 7.63 (m, 1H), 2.88 (d, 3H); MS (EI) for $\text{C}_{15}\text{H}_{13}\text{N}_5\text{O}$: 280.2 (MH^+).

Example 4

3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid

- [0288] N-methyl benzamides, where A is $-\text{C}(=\text{O})-$ and E is $-\text{N}(\text{H})\text{CH}_3$ according to formula I, were prepared in analogous fashion as described above starting from 3-Amino-6-bromo-pyrazine-2-carboxylic acid methylamide.
- [0289] Methyl 3-[5-amino-6-(methylamino)carbonyl] pyrazin-2-yl}benzoate: To a stirred mixture of 3-amino-6-bromo-pyrazine-2-carboxylic acid methylamide (2.29 g, 9.91 mmol, 1.00 equiv.), 3-methoxycarbonylphenylboronic acid (Digital Specialty Chemicals, Inc., 2.68 g, 14.9 mmol, 1.50 equiv.), triethylamine (2.90 g, 28.6 mmol, 2.88 equiv.) in 30 mL of DMF was added [1,1'-bis(diphenylphosphino)-ferrocene]dichloropalladium(II) complex with dichloromethane (1:1), (Aldrich Chemical Company, 405 mg, 0.496 mmol, 0.050 equiv.). The stirred mixture was warmed to 80-90 $^\circ\text{C}$ for 24 hr and then concentrated on a rotary evaporator under reduced pressure. The residue was dissolved in 250 mL of EtOAc, washed three times with 10 mL of 1 N aqueous NaOH, and then saturated aqueous NaCl. The organic solution was filtered through celite, and then concentrated on a rotary evaporator under reduced pressure. The crude product was purified by silica gel chromatography, using EtOAc as eluent, to afford the title compound (2.10 g, 74% yield).
- [0290] 3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid: A flask was charged with methyl 3-[5-amino-6-(methylamino)carbonyl] pyrazin-2-yl}benzoate (2.86 g, 10 mmol) and 30 mL of a solution of 2 N aqueous NaOH. The mixture was refluxed

for 15 min. and then allowed to cool to room temperature. The mixture was extracted with ethyl acetate (10mL). The pH of the aqueous phase was adjusted to pH between 2 – 3 with 10% aqueous HCl with stirring. The resulting pale yellow solid product was collected by filtration, washed with water, and dried, to afford 3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid (2.30 g 84.5% yield): ^1H NMR (400 MHz; DMSO- d_6): 8.83 (s, 1H); 8.80 (s, 1H); 8.51 (s, 1H); 8.41 (d, 1H); 7.93 (d, 1H); 7.65 (br s, 2H); 7.58 (m, 1H); 2.84 (s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{12}\text{N}_4\text{O}_3$: 273 (MH^+).

[0291] 3-amino-6-[3-({[(4-chlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: A flask was charged with 3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid (48.9 mg, 0.18 mmol), 1-hydroxybenzotriazole (HOBT, 39.7 mg, 0.294 mmol), 2.00 mL of tetrahydrofuran, 4-chlorobenzylamine (28.2 mg, 0.2 mmol) and 1-ethyl-(3-dimethylaminopropyl)carbodiimide hydrochloride (66.2 mg, 0.353 mmol). The reaction was stirred at room temperature for 2 h. The reaction was diluted with ethyl acetate, washed sequentially with saturated aqueous NaCl, saturated aqueous sodium carbonate, 1.0 M aqueous hydrochloric acid, and saturated aqueous sodium bicarbonate. The organic layer was dried over sodium sulfate, filtered, and concentrated on a rotary evaporator under reduced pressure. The crude product was purified by chromatography on silica gel using EtOAc as the eluent. After concentration, the product was triturated in 5% ethyl acetate in hexanes. The solid product was filtered, and after drying afforded 3-amino-6-[3-({[(4-chlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide (39.8 mg, 50% yield). ^1H NMR (400 MHz; DMSO- d_6): 9.15 (m, 1H); 8.87 (s, 1H); 8.80 (m, 1H); 8.50 (s, 1H); 8.35 (d, 1H); 7.85 (d, 1H); 7.53 (d, 1H); 7.37 (m, 2H); 4.51 (d, 2H); 2.83 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{Cl}$: 396 (MH^+).

[0292] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0293] 3-amino-6-[(2-oxo-2-phenyl-ethylcarbamoyl)-phenyl]-pyrazine-2-carboxylic acid methylamide: ^1H NMR (400 MHz; DMSO- d_6): 9.13 (br s, 1H); 8.98 (br s, 1H); 8.90 (s, 1H); 8.65 (s, 1H); 8.38 (d, 1H); 8.07 (m, 2H); 7.84 (d, 1H); 7.64 (m, 1H); 6.60 (m, 3H); 4.81 (d, 2H); 2.83 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_3$: 390 (MH^+).

[0294] 3-amino-N-methyl-6-{3-[(4-methylpiperazin-1-yl)carbonyl]phenyl}pyrazine-2-carboxamide: MS (EI) for $\text{C}_{18}\text{H}_{22}\text{N}_6\text{O}_2$: 355 (MH^+).

- [0295] 3-amino-N-methyl-6-(3-([(2-phenylethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{21}N_5O_2$: 376 (MH^+).
- [0296] 3-amino-N-methyl-6-(3-([(4-(methyloxy)phenyl)methyl]amino)carbonyl)phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{21}N_5O_3$: 392 (MH^+).
- [0297] 3-amino-6-(3-([(1R)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{22}H_{21}N_5O_2$: 388 (MH^+).
- [0298] 3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{22}H_{21}N_5O_2$: 388 (MH^+).
- [0299] 3-amino-6-(3-([(4-chlorophenyl)methyl]amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{20}H_{18}N_5O_2Cl$: 396 (MH^+).
- [0300] 3-amino-N-methyl-6-(3-([methyl(phenylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{21}N_5O_2$: 376 (MH^+).
- [0301] 3-amino-6-(3-([(2,4-dichlorophenyl)methyl]amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{20}H_{17}N_5O_2Cl_2$: 431 (MH^+).
- [0302] 3-amino-N-methyl-6-(3-([(4-methylphenyl)methyl]amino)carbonyl)phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{21}N_5O_2$: 376 (MH^+).
- [0303] 3-amino-N-methyl-6-(3-([(4-(trifluoromethyl)phenyl)methyl]amino)carbonyl)phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{18}N_5O_2F_3$: 430 (MH^+).
- [0304] 3-amino-6-(3-([(4-fluorophenyl)methyl]amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{20}H_{18}N_5O_2F$: 380 (MH^+).
- [0305] 3-amino-6-(3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 8.84 (s, 1H); 8.82 (m, 1H); 8.73(d, 1H); 8.47 (m, 1H); 8.31 (d, 1H); 7.82 (d, 1H); 7.52 (t, 1H); 7.12-7.22 (m, 3H); 4.76 (m, 1H); 3.20 (m, 2H); 3.0 (m, 2H) 2.82 (d, 3H); MS (EI) for $C_{22}H_{21}N_5O_2$: 388 (MH^+).
- [0306] 3-amino-N-methyl-6-(3-([(3-(methyloxy)phenyl)methyl]amino)carbonyl)phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{21}N_5O_3$: 392 (MH^+).
- [0307] 3-amino-N-methyl-6-(3-([(1R)-1-[4-(methyloxy)phenyl]ethyl]amino)carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 8.85 (s, 1H); 8.82 (m,

2H); 8.49(s, 1H); 8.38 (d, 1H); 7.80 (d, 1H); 7.65 (br s, 2H); 7.57 (t, 1H); 7.30 (d, 2H); 6.87 (d, 2H); 5.18 (m, 1H); 3.70 (m, 3H); 2.86 (s, 3H) 1.50 (d, 3H); MS (EI) for $C_{22}H_{23}N_5O_3$: 406 (MH^+).

[0308] 3-amino-N-methyl-6-{3-[[[2-(trifluoromethyl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide: MS (EI) for $C_{21}H_{18}N_5O_2F_3$: 430 (MH^+).

[0309] 3-amino-6-[3-[[[3-fluorophenyl]methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{20}H_{18}N_5O_2F$: 380 (MH^+)

[0310] 3-amino-6-[3-[[[1-(4-fluorophenyl)ethyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{21}H_{20}N_5O_2F$: 394 (MH^+)

[0311] 3-amino-6-{3-[[[2,3-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{24}H_{25}N_3O_4$: 420 (MH^+)

[0312] 3-amino-6-(3-{[bis(phenylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{27}H_{25}N_5O_2$: 452 (MH^+)

[0313] 3-amino-6-{3-[[[2,4-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{22}H_{23}N_5O_4$: 422 (MH^+)

[0314] 3-amino-6-(3-{[ethyl(pyridin-4-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{21}H_{22}N_6O_2$: 391 (MH^+)

[0315] 3-amino-N-methyl-6-(3-{[4-(phenylmethyl)piperazin-1-yl]carbonyl}phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{24}H_{26}N_6O_2$: 431 (MH^+)

[0316] 3-amino-N-methyl-6-[3-([1-(phenylmethyl)piperidin-4-yl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 8.84 (s, 1H); 8.82 (m, 1H); 8.42(s, 1H); 8.36 (br s, 1H); 7.80 (m, 1H); 7.65 (br s, 2H); 7.45 (t, 1H); 7.30 (m, 3H); 3.23 (m, 1H); 3.47 (s, 2H); 2.85 (s, 3H) 1.84-2.04 (m, 8H); MS (EI) for $C_{25}H_{28}N_6O_2$: 445 (MH^+)

[0317] 3-amino-N-methyl-6-[3-([5-methylpyrazin-2-yl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: MS (EI) for $C_{19}H_{19}N_7O_2$: 378 (MH^+)

[0318] 3-amino-N-methyl-6-(3-{[(pyridin-2-ylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{19}H_{18}N_6O_2$: 363 (MH^+)

[0319] 3-amino-N-methyl-6-(3-{[(pyridin-3-ylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: MS (EI) for $C_{19}H_{18}N_6O_2$: 363 (MH^+)

- [0320] 3-amino-N-methyl-6-(3-({[(pyridin-4-ylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: : ^1H NMR (400 MHz; DMSO- d_6): 9.21 (s, 1H); 8.89 (s, 1H); 8.26(s, 1H); 8.50 (m, 3H); 8.39 (d, 1H); 7.86 (d, 1H); 7.57 (t, 1H); 7.33 (s, 1H); 4.55 (s, 2H); 2.85 (s, 3H); MS (EI) for $\text{C}_{19}\text{H}_{18}\text{N}_6\text{O}_2$: 363 (MH^+)
- [0321] 3-amino-6-[3-({[(3-chlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{Cl}$: 396 (MH^+)
- [0322] 3-amino-6-(3-({[(cyclohexylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{20}\text{H}_{25}\text{N}_5\text{O}_2$: 368 (MH^+)
- [0323] 3-amino-N-methyl-6-(3-({[(2-phenylpropyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390 (MH^+)
- [0324] 3-amino-6-[3-({[(2-chlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{Cl}$: 396 (MH^+)
- [0325] 3-amino-6-[3-({[2-(2-chlorophenyl)ethyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{O}_2\text{Cl}$: 410 (MH^+)
- [0326] 3-amino-N-methyl-6-{3-[(2-{2-(methyloxy)phenyl}ethyl)amino]carbonyl}phenyl}pyrazine-2-carboxamide: MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_3$: 406 (MH^+)
- [0327] 3-amino-6-[3-({[2-(4-fluorophenyl)ethyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{O}_2\text{F}$: 394 (MH^+)
- [0328] 3-amino-N-methyl-6-[3-({[2-(2-thienyl)ethyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: MS (EI) for $\text{C}_{19}\text{H}_{19}\text{N}_5\text{O}_2\text{S}$: 382 (MH^+)
- [0329] 3-amino-6-{3-[(cyclohexylamino)carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{19}\text{H}_{23}\text{N}_5\text{O}_2$: 354 (MH^+)
- [0330] 1,1-dimethylethyl 4-({[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbonyl]amino}methyl)piperidine-1-carboxylate: MS (EI) for $\text{C}_{24}\text{H}_{32}\text{N}_6\text{O}_4$: 469 (MH^+)
- [0331] 3-amino-N-methyl-6-(3-({[(piperidin-4-ylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: MS (EI) for $\text{C}_{19}\text{H}_{24}\text{N}_6\text{O}_2$: 369 (MH^+)
- [0332] 3-amino-N-methyl-6-[3-({[(3S)-1-(phenylmethyl)pyrrolidin-3-yl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- d_6):

8.62 (s, 1H); 8.35 (s, 1H); 8.03 (br s, 1H); 7.98 (d, 1H); 7.52 (t, 1H); 7.3 (m, 5H); 6.85 (d, 1H); 4.74 (m, 1H); 3.65 (d, 2H); 2.8-3.1 (m, 4H); 2.8 (m, 1H); 2.65 (m, 1H); 2.4 (m, 1H); 2.3 (m, 1H); 1.8 (m, 1H); MS (EI) for $C_{24}H_{26}N_6O_2$: 431 (MH^+)

[0333] 3-amino-N-methyl-6-[3-({[(3R)-1-(phenylmethyl)pyrrolidin-3-yl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: MS (EI) for $C_{24}H_{26}N_6O_2$: 431 (MH^+)

[0334] 3-amino-N-methyl-6-[3-({[2-(phenyloxy)ethyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 8.62 (s, 1H); 8.35 (m, 1H); 7.98 (m, 2H); 7.72 (d, 1H); 7.54 (t, 1H); 7.3 (m, 1H); 6.92 (m, 2H); 6.75 (m, 1H); 4.2 (m, 2H); 3.97 (m, 2H); 3.0 (s, 3H); MS (EI) for $C_{21}H_{21}N_5O_3$: 392 (MH^+)

[0335] 3-amino-N-methyl-6-[3-({[2-[4-(phenylmethyl)piperazin-1-yl]ethyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: MS (EI) for $C_{26}H_{31}N_7O_3$: 473 (MH^+)

[0336] 3-amino-N-methyl-6-[3-({[1-(phenylmethyl)piperidin-4-yl]methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 8.62 (s, 1H); 8.35 (m, 1H); 7.98 (m, 2H); 7.68 (d, 1H); 7.54 (t, 1H); 7.3 (m, 5H); 6.25 (m, 1H); 3.5 (s, 2H); 3.4 (t, 2H); 3.05 (d, 3H); 2.92 (m, 2H); 1.98 (m, 2H); 1.78 (m, 2H); 1.38 (m, 2H); MS (EI) for $C_{26}H_{30}N_6O_2$: 459 (MH^+)

[0337] 3-amino-6-(3-({[(furan-3-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 8.95 (m, 1H); 8.90 (s, 1H); 8.84 (m, 1H); 8.48 (s, 1H); 8.35 (d, 1H); 7.83 (d, 1H); 7.6 (m, 3H); 6.5 (s, 1H); 4.38 (d, 2H); 3.05 (d, 3H); 2.85 (d, 3H); MS (EI) for $C_{18}H_{17}N_5O_3$: 352 (MH^+)

[0338] 3-amino-6-(3-({[(2,3-dihydro-1,4-benzodioxin-2-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{22}H_{21}N_5O_4$: 420 (MH^+)

[0339] 3-amino-6-(3-({[(furan-2-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 9.05 (t, 1H); 8.88 (s, 1H); 8.82 (m, 1H); 8.48 (s, 1H); 8.38 (d, 1H); 7.85 (d, 1H); 7.55 (m, 2H); 6.39 (m, 1H); 6.35 (d, 1H); 3.5 (d, 2H); 2.85 (d, 3H); MS (EI) for $C_{18}H_{17}N_5O_3$: 352 (MH^+)

[0340] 3-amino-N-methyl-6-[3-({[4-(phenylmethyl)morpholin-2-yl]methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: MS (EI) for $C_{25}H_{28}N_6O_3$: 461 (MH^+)

- [0341] 3-amino-6-(3-((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{22}H_{21}N_5O_2$: 420 (MH^+)
- [0342] 3-amino-6-(3-((2,3-dihydro-1-benzofuran-5-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- D_6): 9.07 (t, 1H); 8.88 (s, 1H); 8.85 (m, 1H); 8.52 (s, 1H); 8.38 (d, 1H); 7.88 (d, 1H); 7.58 (t, 1H); 7.22 (s, 1H); 7.08 (d, 1H); 6.7 (d, 1H); 4.5 (m, 2H); 3.2 (m, 2H); 2.85 (d, 3H); MS (EI) for $C_{22}H_{21}N_5O_3$: 404 (MH^+)
- [0343] 3-amino-6-[3-(((6-fluoro-4H-1,3-benzodioxin-8-yl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine : MS (EI) for $C_{22}H_{20}N_5O_4F$: 438 (MH^+)
- [0344] 3-amino-6-{3-[[[3,5-bis(methyloxy)phenyl]methyl]amino)carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- D_6): 9.1 (t, 1H); 8.91 (s, 1H); 8.82(m, 1H); 8.52 (m, 1H); 8.39 (d, 1H); 7.88 (d, 1H); 7.58(t, 1H); 6.52 (m, 2H); 6.38 (m, 1H); 4.48 (d, 2H); 3.76 (s, 6H); 2.85 (d, 3H); MS (EI) for $C_{22}H_{23}N_5O_4$: 422 (MH^+)
- [0345] 3-amino-N-methyl-6-{3-[[[4-(methylsulfonyl)phenyl]methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- D_6): 9.39 (t, 1H); 8.31 (s, 1H); 8.72 (m, 1H); 8.52 (s, 1H); 8.39 (d, 1H); 7.88-7.6 (m, 8H); 4.6 (d, 2H); 3.2 (s, 3H); 2.85 (d, 3H); MS (EI) for $C_{21}H_{21}N_5O_4S$: 440 (MH^+)
- [0346] 3-amino-6-[3-(((5-bromo-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- D_6): 9.19 (t, 1H); 8.84 (s, 1H); 8.82 (m, 1H); 8.52 (m, 1H); 8.38 (d, 1H); 7.86 (d, 1H); 7.54 (m, 2H); 7.5 (m, 3H); 7.2 (t, 1H); 4.54 (d, 2H); 2.84 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2FBr$: 460 (MH^+)
- [0347] 3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- D_6): 9.29 (t, 1H); 8.91 (s, 1H); 8.84 (m, 1H); 8.52 (s, 1H); 8.39 (d, 1H); 7.88 (d, 1H); 7.7 (d, 1H); 7.58 (t, 1H); 7.4 (m, 1H); 7.35 (t, 1H); 4.48 (d, 2H); 2.85 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2FBr$: 458 (MH^+)
- [0348] 3-amino-N-methyl-6-{3-[[[2,4,6-tris(methyloxy)phenyl]methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- D_6): 8.88 (s, 1H); 8.84 (m, 1H); 8.42(s, 1H); 8.3 (d, 1H); 8.18 (m, 1H); 7.88 (d, 1H); 7.5 (t, 1H); 6.2 (s, 2H); 4.4 (d, 2H); 3.8 (s, 9H); 2.85(d, 3H); MS (EI) for $C_{23}H_{25}N_5O_5$: 452 (MH^+)

- [0349] 3-amino-N-methyl-6-[3-({[(3-methylphenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 9.1 (t, 1H); 8.91 (s, 1H); 8.84 (m, 1H); 8.54 (s, 1H); 8.39 (d, 1H); 7.88 (d, 1H); 7.59 (t, 1H); 7.26-7.05 (m, 4H); 4.5 (d, 2H); 2.85 (d, 3H); 2.35 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376 (MH^+)
- [0350] 3-amino-6-[3-({[(4-bromo-2-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 9.18 (t, 1H); 8.92 (s, 1H); 8.84 (m, 1H); 8.56 (s, 1H); 8.39 (d, 1H); 7.88 (d, 1H); 7.59 (m, 2H); 7.4 (m, 2H); 4.5 (d, 2H); 2.85 (t, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{FBr}$: 460 (MH^+)
- [0351] 3-amino-6-[3-({[(2,5-dimethylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.98 (t, 1H); 8.91 (s, 1H); 8.84 (m, 1H); 8.54 (s, 1H); 8.39 (d, 1H); 7.88 (d, 1H); 7.59 (t, 1H); 7.5 (m, 2H); 7.18 (d, 1H); 4.5 (d, 2H); 2.85 (d, 3H); 2.3 (s, 3H); 2.2 (s, 3H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390 (MH^+)
- [0352] 3-amino-6-[3-({[(3,5-bis(trifluoromethyl)phenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 9.3 (t, 1H); 8.91 (s, 1H); 8.84 (m, 1H); 8.54 (s, 1H); 8.39 (d, 1H); 8.05 (m, 3H); 7.85 (d, 1H); 7.6 (t, 2H); 4.7 (d, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{17}\text{N}_5\text{O}_2\text{F}_6$: 498 (MH^+)
- [0353] 3-amino-6-[3-({[(2,2-diphenylethyl)amino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.2 (s, 1H); 8.15 (t, 1H); 8.64 (m, 1H); 8.34 (m, 2H); 7.7 (d, 1H); 7.5 (t, 1H); 7.39-7.21 (m, 10H); 7.2 (m, 2H); 4.46 (t, 1H); 3.95 (m, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{27}\text{H}_{25}\text{N}_5\text{O}_2$: 452 (MH^+)
- [0354] 3-amino-6-[3-({[(4-ethylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: (400 MHz; DMSO- D_6): 9.8 (t, 1H); 8.88 (s, 1H); 8.82 (m, 1H); 8.5 (s, 1H); 8.36 (d, 1H); 7.86 (d, 1H); 7.56 (t, 2H); 7.25 – 7.14 (m, 4H); 4.4 (d, 2H); 2.85 (d, 3H); 2.6 (m, 2H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390 (MH^+)
- [0355] 3-amino-N-methyl-6-[3-({[(4-propylphenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: (400 MHz; DMSO- D_6): 9.08 (t, 1H); 8.88 (s, 1H); 8.81 (m, 1H); 8.5 (s, 1H); 8.38 (d, 1H); 7.86 (d, 1H); 7.55 (t, 1H); 7.18 (m, 4H); 4.5 (d, 2H); 2.85 (d, 3H); 2.55 (m, 2H); 1.5 (m, 2H); 1.08 (t, 3H); MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_5\text{O}_2$: 404 (MH^+).

- [0356] 3-amino-6-(3-([(1,1-dimethylethyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.92 (s, 1H), 8.88 (m, 1H), 8.38 (m, 2H), 7.90 (m, 1H), 7.81 (m, 1H), 7.71 (br s, 2H), 7.53 (m, 1H), 2.87 (d, 3H), 1.43 (s, 9H); MS (EI) for $\text{C}_{17}\text{H}_{21}\text{N}_5\text{O}_2$: 328.5 (MH^+).
- [0357] 3-amino-6-[3-([(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.97 (m, 1H), 8.87 (m, 1H), 8.62 (m, 2H), 8.41 (m, 1H), 7.96 (m, 1H), 7.74 (br s, 2H), 7.59 (m, 1H), 7.26 (m, 4H), 5.52 (m, 1H), 5.14 (m, 1H), 4.55 (m, 1H), 3.14 (m, 1H), 2.93 (m, 1H), 2.85 (m, 3H); MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_5\text{O}_3$: 404.5 (MH^+).
- [0358] 3-amino-6-[3-([(1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.97 (m, 1H), 8.87 (m, 1H), 8.62 (m, 2H), 8.41 (m, 1H), 7.96 (m, 1H), 7.74 (br s, 2H), 7.59 (m, 1H), 7.26 (m, 4H), 5.52 (m, 1H), 5.14 (m, 1H), 4.55 (m, 1H), 3.14 (m, 1H), 2.93 (m, 1H), 2.85 (m, 3H); MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_5\text{O}_3$: 404.5 (MH^+).
- [0359] 3-amino-N-methyl-6-[3-[(phenylamino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.34 (m, 1H), 8.97 (s, 1H), 8.90 (m, 1H), 8.61 (m, 1H), 8.42 (m, 1H), 7.93 (m, 1H), 7.81 (m, 2H), 7.70 (br s, 2H), 7.63 (m, 1H), 7.38 (m, 2H), 7.13 (m, 1H), 2.87 (m, 3H); MS (EI) for $\text{C}_{19}\text{H}_{17}\text{N}_5\text{O}_2$: 348.4 (MH^+).
- [0360] 3-amino-6-[3-(1,3-dihydro-2H-isoindol-2-ylcarbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.92 (s, 1H), 8.86 (m, 1H), 8.33 (m, 2H), 7.67 (br s, 2H), 7.56 (m, 2H), 7.43 (m, 1H), 7.29 (m, 3H), 4.91 (s, 2H), 4.79 (s, 2H), 2.84 (m, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_2$: 374.5 (MH^+).
- [0361] 3-amino-N-methyl-6-[3-([(1R)-1-phenylethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.86 (m, 2H), 8.48 (s, 1H), 8.36 (m, 1H), 7.84 (m, 1H), 7.71 (br s, 2H), 7.54 (m, 1H), 7.40 (m, 2H), 7.32 (m, 2H), 7.21 (s, 1H), 5.20 (m, 1H), 2.85 (s, 3H), 1.53 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376.5 (MH^+).
- [0362] 3-amino-N-methyl-6-[3-([(1S)-1-phenylethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.86 (m, 2H), 8.48 (s, 1H), 8.36 (m, 1H), 7.84 (m, 1H), 7.71 (br s, 2H), 7.54 (m, 1H), 7.40 (m, 2H), 7.32 (m, 2H), 7.21 (s, 1H), 5.20 (m, 1H), 2.85 (s, 3H), 1.53 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376.5 (MH^+).

- [0363] 3-amino-N-methyl-6-[3-({[(1R)-1-phenylpropyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.83 (m, 2H), 8.47 (s, 1H), 8.36 (m, 1H), 7.83 (m, 1H), 7.71 (br s, 2H), 7.54 (t, 1H), 7.39 (m, 2H), 7.32 (m, 2H), 7.21 (m, 1H), 4.94 (m, 1H), 2.85 (s, 3H), 1.86 (m, 2H), 0.94 (m, 3H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390.5 (MH^+).
- [0364] 3-amino-N-methyl-6-[3-({[(1S)-1-phenylpropyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.83 (m, 2H), 8.47 (s, 1H), 8.36 (m, 1H), 7.83 (m, 1H), 7.71 (br s, 2H), 7.54 (t, 1H), 7.39 (m, 2H), 7.32 (m, 2H), 7.21 (m, 1H), 4.94 (m, 1H), 2.85 (s, 3H), 1.86 (m, 2H), 0.94 (m, 3H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390.5 (MH^+).
- [0365] 3-amino-N-methyl-6-[3-({[(1R,2S)-2-phenylcyclopropyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.89 (s, 1H), 8.83 (m, 1H), 8.76 (d, 1H), 8.44 (s, 1H), 8.34 (d, 1H), 7.81 (d, 1H), 7.72 (br s, 2H), 7.54 (t, 1H), 7.27 (m, 2H), 7.17 (m, 3H), 3.06 (m, 1H), 2.85 (d, 3H), 2.12 (m, 1H), 1.39 (m, 1H), 1.26 (m, 1H); MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_5\text{O}_2$: 388.5 (MH^+).
- [0366] 3-amino-N-methyl-6-(3-({[(1R)-1,2,3,4-tetrahydronaphthalen-1-ylamino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.88 (s, 1H), 8.84 (m, 2H), 8.50 (s, 1H), 8.38 (d, 1H), 7.90 (d, 1H), 7.71 (brs, 2H), 7.56 (t, 1H), 7.22 (m, 1H), 7.16 (m, 3H), 5.29 (m, 1H), 2.84 (d, 3H), 2.79 (m, 2H), 2.02 (m, 2H), 1.84 (m, 2H); MS (EI) for $\text{C}_{23}\text{H}_{23}\text{N}_5\text{O}_2$: 402.5 (MH^+).
- [0367] 3-amino-N-methyl-6-(3-({[(1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.88 (s, 1H), 8.84 (m, 2H), 8.50 (s, 1H), 8.38 (d, 1H), 7.90 (d, 1H), 7.71 (brs, 2H), 7.56 (t, 1H), 7.22 (m, 1H), 7.16 (m, 3H), 5.29 (m, 1H), 2.84 (d, 3H), 2.79 (m, 2H), 2.02 (m, 2H), 1.84 (m, 2H); MS (EI) for $\text{C}_{23}\text{H}_{23}\text{N}_5\text{O}_2$: 402.5 (MH^+).
- [0368] 3-amino-N-methyl-6-[3-({[3,4,5-tris(methyloxy)phenyl]methyl}amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.05 (s, 1H), 8.89 (s, 1H), 8.82 (m, 1H), 8.50 (s, 1H), 8.36 (d, 1H), 7.87 (d, 1H), 7.71 (br s, 2H), 7.56 (m, 1H), 6.67 (s, 2H), 4.45 (s, 2H), 3.76 (s, 6H), 3.62 (s, 3H), 2.84 (s, 3H); MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_5\text{O}_5$: 452.5 (MH^+).
- [0369] 3-amino-N-methyl-6-[3-({[(1R,2R)-2-[(phenylmethyl)oxy]cyclopentyl]amino}carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.89 (s,

1H), 8.82 (m, 1H), 8.42 (m, 2H), 8.34 (d, 1H), 7.81 (m, 1H), 7.75 (br s, 2H), 7.55 (m, 1H), 7.30 (m, 4H), 7.23 (m, 1H), 4.57 (dd, 2H), 4.32 (m, 1H), 3.94 (m, 1H), 2.83 (d, 3H), 2.07 (m, 1H), 1.95 (m, 1H), 1.70 (m, 4H); MS (EI) for $C_{25}H_{27}N_5O_3$: 446.6 (MH^+).

[0370] 3-amino-N-methyl-6-{3-[(1S,2S)-2-[(phenylmethyl)oxy]cyclopentyl]amino)carbonyl}phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.89 (s, 1H), 8.82 (m, 1H), 8.42 (m, 2H), 8.34 (d, 1H), 7.81 (m, 1H), 7.75 (br s, 2H), 7.55 (m, 1H), 7.30 (m, 4H), 7.23 (m, 1H), 4.57 (dd, 2H), 4.32 (m, 1H), 3.94 (m, 1H), 2.83 (d, 3H), 2.07 (m, 1H), 1.95 (m, 1H), 1.70 (m, 4H); MS (EI) for $C_{25}H_{27}N_5O_3$: 446.5 (MH^+).

[0371] 3-amino-6-(3-[(biphenyl-2-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.02 (t, 1H), 8.90 (s, 1H), 8.84 (m, 1H), 8.49 (s, 1H), 8.38 (d, 1H), 7.86 (d, 1H), 7.74 (br s, 2H), 7.56 (t, 1H), 7.47 (m, 5H), 7.38 (m, 3H), 7.26 (m, 1H), 4.49 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{26}H_{23}N_5O_2$: 438.5 (MH^+).

[0372] 3-amino-6-(3-[(biphenyl-4-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.19 (t, 1H), 8.92 (s, 1H), 8.84 (m, 1H), 8.55 (s, 1H), 8.40 (d, 1H), 7.91 (d, 1H), 7.75 (br s, 2H), 7.65 (m, 4H), 7.59 (t, 1H), 7.46 (m, 4H), 7.35 (m, 1H), 4.58 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{26}H_{23}N_5O_2$: 438.5 (MH^+).

[0373] 3-amino-N-methyl-6-{3-[[[4-(phenyloxy)phenyl]methyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.14 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.54 (s, 1H), 8.39 (d, 1H), 7.89 (d, 1H), 7.75 (br s, 2H), 7.58 (t, 1H), 7.38 (m, 4H), 7.12 (m, 1H), 6.99 (m, 4H), 4.53 (d, 2H), 2.87 (d, 3H); MS (EI) for $C_{26}H_{23}N_5O_3$: 454.5 (MH^+).

[0374] 3-amino-N-methyl-6-{3-[[[3-(phenyloxy)phenyl]methyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.14 (t, 1H), 8.90 (s, 1H), 8.83 (m, 1H), 8.51 (s, 1H), 8.39 (d, 1H), 7.86 (d, 1H), 7.74 (br s, 2H), 7.57 (t, 1H), 7.37 (m, 3H), 7.13 (m, 2H), 7.01 (m, 3H), 6.88 (m, 1H), 4.52 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{26}H_{23}N_5O_3$: 454.6 (MH^+).

[0375] 3-amino-6-(3-[(biphenyl-3-ylmethyl)amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.19 (t, 1H), 8.92 (s, 1H), 8.84 (m, 1H), 8.56 (s, 1H), 8.39 (d, 1H), 7.90 (d, 1H), 7.69 (br s, 2H), 7.65 (m, 3H), 7.57 (m, 2H),

7.46 (m, 3H), 7.38 (m, 2H), 4.63 (d, 2H), 2.83 (d, 3H); MS (EI) for $C_{26}H_{23}N_5O_2$: 438.4 (MH^+).

[0376] 3-amino-N-methyl-6-(3-([(1-methyl-1-phenylethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.96 (s, 1H), 8.87 (m, 1H), 8.54 (s, 1H), 8.48 (s, 1H), 8.38 (d, 1H), 7.82 (d, 1H), 7.75 (br s, 2H), 7.55 (t, 1H), 7.42 (m, 2H), 7.30 (m, 2H), 7.18 (m, 1H), 2.85 (d, 3H), 1.71 (s, 6H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390.4 (MH^+).

[0377] 3-amino-6-[3-([(2,4-difluorophenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.13 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.52 (s, 1H), 8.40 (d, 1H), 7.87 (d, 1H), 7.73 (br s, 2H), 7.58 (t, 1H), 7.48 (m, 1H), 7.25 (m, 1H), 7.08 (m, 1H), 4.53 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2F_2$: 398.3 (MH^+).

[0378] 3-amino-6-[3-([(2-chloro-6-(phenyloxy)phenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.85 (s, 1H), 8.83 (m, 1H), 8.69 (t, 1H), 8.36 (s, 1H), 8.33 (d, 1H), 7.77 (d, 1H), 7.72 (br s, 2H), 7.51 (t, 1H), 7.34 (m, 4H), 7.05 (m, 3H), 6.85 (d, 1H), 4.67 (d, 2H), 2.83 (d, 3H); MS (EI) for $C_{26}H_{22}N_5O_3Cl$: 488.4 (MH^+).

[0379] 3-amino-6-[3-([(2,5-dichlorophenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.18 (t, 1H), 8.92 (s, 1H), 8.84 (m, 1H), 8.56 (s, 1H), 8.41 (d, 1H), 7.91 (d, 1H), 7.74 (br s, 2H), 7.60 (t, 1H), 7.54 (d, 1H), 7.41 (m, 2H), 4.58 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2Cl_2$: 430.4 (MH^+).

[0380] 3-amino-6-[3-([(2,6-bis(methyloxy)phenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.90 (s, 1H), 8.86 (m, 1H), 8.45 (s, 1H), 8.33 (d, 1H), 8.28 (t, 1H), 7.84 (d, 1H), 7.71 (br s, 2H), 7.53 (t, 1H), 7.27 (t, 1H), 6.69 (d, 2H), 4.50 (d, 2H), 3.80 (s, 6H), 2.85 (d, 3H); MS (EI) for $C_{22}H_{23}N_5O_4$: 422.4 (MH^+).

[0381] 3-amino-6-[3-([(2-chloro-6-methylphenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.89 (s, 1H), 8.84 (m, 1H), 8.69 (m, 1H), 8.47 (s, 1H), 8.36 (d, 1H), 7.85 (d, 1H), 7.69 (br s, 2H), 7.54 (t, 1H), 7.32 (m, 1H), 7.24 (m, 2H), 4.65 (d, 2H), 2.85 (d, 3H), 2.45 (s, 3H); MS (EI) for $C_{21}H_{20}N_5O_2Cl$: 410.4 (MH^+).

- [0382] 3-amino-N-methyl-6-[3-({[(2,4,5-trifluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.14 (t, 1H), 8.92 (s, 1H), 8.85 (m, 1H), 8.53 (s, 1H), 8.40 (d, 1H), 7.87 (d, 1H), 7.75 (br s, 2H), 7.58 (m, 2H), 7.50 (m, 1H), 4.52 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{16}\text{N}_5\text{O}_2\text{F}_3$: 416.4 (MH^+).
- [0383] 3-amino-6-[3-({[(2,3-dichlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.22 (t, 1H), 8.93 (s, 1H), 8.86 (m, 1H), 8.57 (s, 1H), 8.42 (d, 1H), 7.90 (d, 1H), 7.76 (br s, 2H), 7.60 (m, 2H), 7.39 (m, 2H), 4.62 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{Cl}_2$: 430.32 (MH^+).
- [0384] 3-amino-6-[3-({[(4-chloro-2-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.16 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.53 (s, 1H), 8.40 (d, 1H), 7.87 (d, 1H), 7.80 (br s, 2H), 7.58 (t, 1H), 7.45 (m, 2H), 7.29 (d, 1H), 4.53 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{FCl}$: 414.4 (MH^+).
- [0385] 3-amino-6-[3-({[(3,5-dichlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.21 (t, 1H), 8.92 (s, 1H), 8.85 (m, 1H), 8.54 (s, 1H), 8.40 (d, 1H), 7.88 (d, 1H), 7.73 (br s, 2H), 7.59 (t, 1H), 7.52 (m, 1H), 7.42 (m, 2H), 4.54 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{Cl}_2$: 430.3 (MH^+).
- [0386] 3-amino-6-[3-({[(5-chloro-2-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.17 (t, 1H), 8.92 (s, 1H), 8.86 (m, 1H), 8.54 (s, 1H), 8.41 (d, 1H), 7.89 (d, 1H), 7.77 (br s, 2H), 7.59 (t, 1H), 7.46 (m, 1H), 7.41 (m, 1H), 7.28 (m, 1H), 4.55 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{FCl}$: 414.4 (MH^+).
- [0387] 3-amino-6-[3-({[(3,4-dichlorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.21 (t, 1H), 8.91 (s, 1H), 8.85 (m, 1H), 8.53 (s, 1H), 8.40 (d, 1H), 7.88 (d, 1H), 7.72 (br s, 2H), 7.60 (m, 3H), 7.36 (d, 1H), 4.53 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{Cl}_2$: 430.3 (MH^+).
- [0388] 3-amino-6-[3-({[(4'-fluorobiphenyl-2-yl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.03 (t, 1H), 8.90 (s, 1H), 8.84 (m, 1H), 8.49 (s, 1H), 8.38 (d, 1H), 7.85 (d, 1H), 7.71 (br s, 2H), 7.56 (t, 1H), 7.49 (m, 3H), 7.42 – 7.22 (m, 5H), 4.46 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{26}\text{H}_{22}\text{N}_5\text{O}_2\text{F}$: 456.4 (MH^+).

- [0389] 3-amino-6-[3-({[(4-fluoro-3-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): (9.12 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.53 (s, 1H), 8.39 (d, 1H), 7.89 (d, 1H), 7.73 (br s, 2H), 7.58 (t, 1H), 7.26 (m, 1H), 7.20 (m, 1H), 7.10 (m, 1H), 4.48 (d, 2H), 2.85 (d, 3H), 2.23 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{O}_2\text{F}$: 394.4 (MH^+).
- [0390] 3-amino-6-[3-({[(3-chloro-2-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.19 (t, 1H), 8.92 (s, 1H), 8.84 (m, 1H), 8.54 (s, 1H), 8.40 (d, 1H), 7.88 (d, 1H), 7.76 (br s, 2H), 7.58 (t, 1H), 7.51 (t, 1H), 7.41 (t, 1H), 7.22 (t, 1H), 4.60 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{FCl}$: 414.4 (MH^+).
- [0391] 3-amino-6-[3-({[(6-chloro-2-fluoro-3-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.89 (s, 1H), 8.84 (m, 1H), 8.48 (s, 1H), 8.37 (d, 2H), 7.85 (d, 1H), 7.74 (br s, 2H), 7.55 (t, 1H), 7.27 (m, 2H), 4.64 (d, 2H), 2.85 (d, 3H), 2.24 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_2\text{FCl}$: 428.4 (MH^+).
- [0392] 3-amino-6-[3-({[(4-iodophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.17 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.52 (s, 1H), 8.39 (d, 1H), 7.88 (d, 1H), 7.74 (br s, 2H), 7.71 (d, 2H), 7.58 (t, 1H), 7.18 (d, 2H), 4.49 (d, 2H), 2.86 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{I}$: 488.2 (MH^+).
- [0393] 3-amino-6-[3-({[(3-chloro-4-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.16 (t, 1H), 8.92 (s, 1H), 8.85 (m, 1H), 8.53 (s, 1H), 8.39 (d, 1H), 7.88 (d, 1H), 7.75 (br s, 2H), 7.58 (t, 1H), 7.39 (s, 1H), 7.32 (d, 1H), 7.23 (d, 1H), 4.49 (d, 2H), 2.87 (d, 3H), 2.31 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{O}_2\text{Cl}$: 410.4 (MH^+).
- [0394] 3-amino-6-[3-({[(2,5-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.13 (t, 1H), 8.89 (s, 1H), 8.82 (m, 1H), 8.52 (s, 1H), 8.38 (d, 1H), 7.86 (d, 1H), 7.71 (br s, 2H), 7.56 (t, 1H), 7.31 – 7.10 (m, 3H), 4.55 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{F}_2$: 398.5 (MH^+).
- [0395] 3-amino-6-[3-({[(2,6-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.95 (t, 1H), 8.86 (s, 1H), 8.80 (m, 1H), 8.45 (s, 1H), 8.34 (d, 1H), 7.81 (d, 1H), 7.71 (br s, 2H), 7.53 (t, 1H),

7.38 (m, 1H), 7.09 (m, 1H), 4.56 (d, 2H), 2.84 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2F_2$: 398.4 (MH^+).

[0396] 3-amino-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.18 (t, 1H), 8.89 (s, 1H), 8.82 (m, 1H), 8.52 (s, 1H), 8.37 (d, 1H), 7.86 (d, 1H), 7.71 (br s, 2H), 7.56 (t, 1H), 7.15 – 7.01 (m, 3H), 4.54 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2F_2$: 398.5 (MH^+).

[0397] 3-amino-6-[3-(((3-iodophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.14 (t, 1H), 8.88 (s, 1H), 8.82 (m, 1H), 8.50 (s, 1H), 8.37 (d, 1H), 7.86 (d, 1H), 7.71 (s, 1H), 7.67 (br s, 2H), 7.64 – 7.52 (m, 2H), 7.36 (d, 1H), 7.14 (t, 1H), 4.48 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{18}N_5O_2I$: 488.4 (MH^+).

[0398] 3-amino-N-methyl-6-[3-(((2-piperidin-1-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, $CDCl_3$): 8.63 (s, 1H), 8.36 (m, 1H), 8.11 (m, 1H), 7.95 (m, 2H), 7.72 (d, 1H), 7.50 (t, 1H), 7.32 (m, 1H), 7.26 (m, 1H), 7.16 (d, 1H), 7.08 (t, 1H), 4.78 (d, 2H), 3.02 (d, 3H), 2.92 (m, 4H), 1.78 – 1.49 (m, 6H); MS (EI) for $C_{25}H_{28}N_6O_2$: 445.5 (MH^+).

[0399] 3-amino-N-methyl-6-[3-(((2-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.04 (t, 1H), 8.90 (s, 1H), 8.82 (m, 1H), 8.54 (m, 1H), 8.38 (d, 1H), 7.88 (d, 1H), 7.70 (br s, 2H), 7.57 (t, 1H), 7.28 (m, 3H), 7.15 (m, 1H), 4.52 (d, 2H), 2.85 (d, 3H), 2.51 (s, 3H); MS (EI) for $C_{21}H_{21}N_5O_2S$: 408.4 (MH^+).

[0400] 3-amino-6-[3-(((2-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.15 (t, 1H), 8.92 (s, 1H), 8.85 (m, 1H), 8.56 (m, 1H), 8.41 (d, 1H), 7.89 (d, 1H), 7.73 (br s, 2H), 7.59 (t, 1H), 7.48 (m, 2H), 7.24 (m, 1H), 4.57 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2FCl$: 414.4 (MH^+).

[0401] 3-amino-6-[3-(((3,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.05 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.52 (m, 1H), 8.37 (d, 1H), 7.88 (d, 1H), 7.74 (br s, 2H), 7.57 (t, 1H), 6.98 (m, 1H), 6.89 (m, 2H), 4.47 (d, 2H), 3.74 (s, 3H), 3.73 (s, 3H), 2.85 (d, 3H); MS (EI) for $C_{22}H_{23}N_5O_4$: 422.5 (MH^+).

- [0402] 3-amino-N-methyl-6-{3-[(phenylmethyl)amino]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.65 (s, 1H), 8.60 (m, 1H), 7.58 (br s, 2H), 7.42 (m, 1H), 7.37 – 7.19 (m, 6H), 7.13 (t, 1H), 6.60 (d, 1H), 6.23 (m, 1H), 4.57 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{19}\text{H}_{19}\text{N}_5\text{O}$: 334.4 (MH^+).
- [0403] 3-amino-6-[3-([[(2-amino-6-fluorophenyl)methyl]amino]carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.01 (t, 1H), 8.88 (s, 1H), 8.82 (m, 1H), 8.48 (s, 1H), 8.36 (d, 1H), 7.86 (d, 1H), 7.71 (br s, 2H), 7.54 (t, 1H), 6.96 (m, 1H), 6.45 (d, 1H), 6.32 (t, 1H), 5.66 (s, 2H), 4.45 (d, 2H), 2.84 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_6\text{O}_2\text{F}$: 395.4 (MH^+).
- [0404] 3-amino-6-{3-([[(4-fluoro-2-(trifluoromethyl)phenyl)methyl]amino]carbonyl)phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.23 (t, 1H), 8.93 (s, 1H), 8.84 (m, 1H), 8.57 (m, 1H), 8.42 (d, 1H), 7.91 (d, 1H), 7.76 (br s, 2H), 7.69 – 7.52 (m, 4H), 4.68 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{17}\text{N}_5\text{O}_2\text{F}_4$: 448.4 (MH^+).
- [0405] 3-amino-N-methyl-6-{3-([[(6-(trifluoromethyl)pyridin-3-yl)methyl]amino]carbonyl)phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.29 (t, 1H), 8.91 (s, 1H), 8.85 (m, 1H), 8.79 (s, 1H), 8.54 (s, 1H), 8.40 (d, 1H), 8.05 (d, 1H), 7.89 (m, 2H), 7.76 (br s, 2H), 7.59 (t, 1H), 4.65 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_6\text{O}_2\text{F}_3$: 431.4 (MH^+).
- [0406] 3-amino-6-(3-{[2-(2,5-difluorophenyl)hydrazino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.52 (s, 1H), 8.93 (s, 1H), 8.84 (m, 1H), 8.59 (s, 1H), 8.42 (d, 1H), 8.20 (s, 1H), 7.88 (d, 1H), 7.74 (br s, 2H), 7.60 (t, 1H), 7.15 (m, 1H), 6.63 (m, 1H), 6.51 (m, 1H), 2.86 (d, 3H); MS (EI) for $\text{C}_{19}\text{H}_{16}\text{N}_6\text{O}_2\text{F}_2$: 399.4 (MH^+).
- [0407] 3-amino-N-methyl-6-(3-{[2-(2,3,5,6-tetrafluoro-4-methylphenyl)hydrazino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.76 (s, 1H), 8.88 (s, 1H), 8.81 (m, 1H), 8.50 (s, 1H), 8.39 (d, 1H), 8.19 (s, 1H), 7.83 (d, 1H), 7.73 (br s, 2H), 7.57 (t, 1H), 2.87 (d, 3H), 2.15 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{16}\text{N}_6\text{O}_2\text{F}_4$: 449.4 (MH^+).
- [0408] 3-amino-6-(3-{[2-(2-fluorophenyl)hydrazino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.50 (s, 1H), 8.92 (s, 1H), 8.84 (m, 1H), 8.58 (s, 1H), 8.41 (d, 1H), 7.88 (m, 2H), 7.74 (br s, 2H), 7.60 (t, 1H), 7.10 (m, 1H),

6.98 (m, 1H), 6.88 (m, 1H), 6.75 (m, 1H), 2.86 (d, 3H); MS (EI) for $C_{19}H_{17}N_6O_2F$: 381.4 (MH^+).

[0409] 3-amino-6-(3-{[2-(2,4-difluorophenyl)hydrazino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 10.51 (s, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.57 (s, 1H), 8.41 (d, 1H), 7.85 (m, 2H), 7.74 (br s, 2H), 7.59 (t, 1H), 7.17 (m, 1H), 6.89 (m, 2H), 2.87 (d, 3H); MS (EI) for $C_{19}H_{16}N_6O_2F_2$: 399.4 (MH^+).

[0410] 3-amino-N-methyl-6-(3-{[2-(2,3,5,6-tetrafluorophenyl)hydrazino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 10.80 (s, 1H), 8.88 (s, 1H), 8.81 (m, 1H), 8.50 (s, 1H), 8.40 (m, 2H), 7.84 (d, 1H), 7.73 (br s, 2H), 7.58 (t, 1H), 7.20 (m, 1H), 2.87 (d, 3H); MS (EI) for $C_{19}H_{14}N_6O_2F_4$: 435.4 (MH^+).

[0411] 3-Amino-6-[3-({[(3,5-dimethylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.08 (t, 1H), 8.91 (s, 1H), 8.84 (q, 1H), 8.54-8.52 (m, 1H), 8.39 (d, 1H), 7.89 (d, 1H), 7.58 (t, 1H), 6.95 (s, 2H), 6.88 (s, 1H), 4.46 (d, 2H), 4.10-3.70 (m, 2H), 2.85 (d, 3H), 2.25 (s, 6H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390 (MH^+).

[0412] 3-amino-6-[3-({[(5-chloro-2-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.06 (t, 1H), 8.91 (s, 1H), 8.84 (q, 1H), 8.56-8.53 (m, 1H), 8.40 (d, 1H), 8.00-7.30 (m, 2H), 7.90 (d, 1H), 7.59 (t, 1H), 7.29 (s, 1H), 7.24-7.22 (m, 2H), 4.50 (d, 2H), 2.86 (d, 3H), 2.34 (s, 3H); MS (EI) for $C_{21}H_{20}N_5O_2Cl$: 410 (MH^+).

[0413] 3-amino-6-[3-({[(2,4-dimethylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.96 (t, 1H), 8.91 (s, 1H), 8.85 (q, 1H), 8.54-8.52 (m, 1H), 8.39 (d, 1H), 7.89 (d, 1H), 7.58 (t, 1H), 7.17 (d, 1H), 7.02-6.95 (m, 2H), 4.46 (d, 2H), 4.00-3.30 (m, 2H), 2.85 (d, 3H), 2.31 (s, 3H), 2.25 (s, 3H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390 (MH^+).

[0414] 3-amino-6-[3-({[(2,3-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.20 (t, 1H), 8.92 (s, 1H), 8.85 (q, 1H), 8.55-8.52 (m, 1H), 8.40 (d, 1H), 7.88 (d, 1H), 7.58 (t, 1H), 7.40-7.30 (m, 1H), 7.28-7.16 (m, 2H), 4.60 (d, 2H), 4.20-3.80 (m, 2H), 2.85 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2F_2$: 398 (MH^+).

[0415] 3-amino-N-methyl-6-[3-({[4-(methylthio)phenyl]methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.13 (t, 1H), 8.91 (s,

1H), 8.84 (q, 1H), 8.53-8.51 (m, 1H), 8.39 (d, 1H), 8.00-7.30 (m, 2H), 7.88 (d, 1H), 7.58 (t, 1H), 7.31 (d, 2H), 7.24 (d, 2H), 4.49 (d, 2H), 2.85 (d, 3H), 2.45 (s, 3H); MS (EI) for $C_{21}H_{21}N_5O_2S$: 408 (MH^+).

[0416] 3-amino-N-methyl-6-(3-{{(2-methylpropyl)amino}carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.84 (q, 1H), 8.56 (t, 1H), 8.48-8.45 (m, 1H), 8.36 (dt, 1H), 8.20-7.30 (m, 2H), 7.83 (dt, 1H), 7.56 (t, 1H), 3.13 (dd, 2H), 2.86 (d, 3H), 1.94-1.82 (m, 1H), 0.92 (d, 6H); MS (EI) for $C_{17}H_{21}N_5O_2$: 328 (MH^+).

[0417] 3-amino-N-methyl-6-(3-{{(3-{{(trifluoromethyl)oxy}phenyl}methyl)amino}carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.24 (t, 1H), 8.91 (s, 1H), 8.85 (q, 1H), 8.55-8.53 (m, 1H), 8.40 (d, 1H), 8.20-7.20 (m, 2H), 7.88 (d, 1H), 7.59 (t, 1H), 7.49 (t, 1H), 7.40 (d, 1H), 7.35 (s, 1H), 7.26 (d, 1H), 4.58 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{21}H_{18}N_5O_3F_3$: 446 (MH^+).

[0418] 3-amino-6-(3-{{(cyclopropylmethyl)amino}carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.85 (q, 1H), 8.67 (t, 1H), 8.49-8.46 (m, 1H), 8.36 (dt, 1H), 8.10-7.20 (m, 2H), 7.85 (dt, 1H), 7.56 (t, 1H), 3.19 (dd, 2H), 2.86 (d, 3H), 1.12-1.01 (m, 1H), 0.49-0.43 (m, 2H), 0.29-0.23 (m, 2H); MS (EI) for $C_{17}H_{19}N_5O_2$: 326 (MH^+).

[0419] 3-amino-6-{3-{{(cyclopentylamino)carbonyl}phenyl}}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.86 (q, 1H), 8.44-8.42 (m, 1H), 8.39-8.33 (m, 2H), 8.10-7.20 (m, 2H), 7.82 (dt, 1H), 7.54 (t, 1H), 4.31-4.21 (m, 1H), 2.85 (d, 3H), 1.99-1.86 (m, 2H), 1.79-1.66 (m, 2H), 1.62-1.50 (m, 4H); MS (EI) for $C_{18}H_{21}N_5O_2$: 340 (MH^+).

[0420] 3-amino-N-methyl-6-(3-{{(4-methylphenyl)amino}carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 10.3 (s, 1H), 8.95 (s, 1H), 8.89 (q, 1H), 8.59-8.57 (m, 1H), 8.40 (d, 1H), 8.00-7.30 (m, 2H), 7.90 (d, 1H), 7.69 (d, 2H), 7.62 (t, 1H), 7.18 (d, 2H), 2.86 (d, 3H), 2.30 (s, 3H); MS (EI) for $C_{20}H_{19}N_5O_2$: 362 (MH^+).

[0421] 3-amino-N-methyl-6-[3-{{(4-(methyloxy)phenyl)amino}carbonyl}phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 10.2 (s, 1H), 8.95 (s, 1H), 8.89 (q, 1H), 8.59-8.57 (m, 1H), 8.40 (dt, 1H), 8.00-7.40 (m, 2H), 7.90 (dt, 1H), 7.73-7.68 (m, 2H), 7.61 (t, 1H), 6.98-6.93 (m, 2H), 3.76 (s, 3H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{19}N_5O_3$: 378 (MH^+).

- [0422] 3-amino-6-(3-([(2,2-dimethylpropyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.92 (s, 1H), 8.85 (q, 1H), 8.49-8.43 (m, 2H), 8.36 (dt, 1H), 8.00-7.40 (m, 2H), 7.83 (dt, 1H), 7.56 (t, 1H), 3.16 (d, 2H), 2.85 (d, 3H), 0.93 (s, 9H); MS (EI) for $\text{C}_{18}\text{H}_{23}\text{N}_5\text{O}_2$: 342 (MH^+).
- [0423] 3-amino-6-(3-([(cyclopentylmethyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.91 (s, 1H), 8.85 (q, 1H), 8.57 (t, 1H), 8.47-8.44 (m, 1H), 8.36 (d, 1H), 8.00-7.30 (m, 2H), 7.82 (d, 1H), 7.55 (t, 1H), 3.26-3.21 (m, 2H), 2.86 (d, 3H), 2.23-2.12 (m, 1H), 1.72-1.45 (m, 6H), 1.33-1.22 (m, 2H); MS (EI) for $\text{C}_{19}\text{H}_{23}\text{N}_5\text{O}_2$: 354 (MH^+).
- [0424] 3-amino-6-{3-[(cyclopropylamino)carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.90 (s, 1H), 8.85 (q, 1H), 8.55-8.52 (m, 1H), 8.43-8.41 (m, 1H), 8.35 (d, 1H), 8.00-7.40 (m, 2H), 7.80 (d, 1H), 7.54 (t, 1H), 2.91-2.84 (m, 4H), 0.77-0.57 (m, 4H); MS (EI) for $\text{C}_{16}\text{H}_{17}\text{N}_5\text{O}_2$: 312 (MH^+).
- [0425] 3-amino-N-methyl-6-[3-([(2,3,4-trifluorophenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.17 (t, 1H), 8.91 (s, 1H), 8.84 (q, 1H), 8.53-8.51 (m, 1H), 8.40 (d, 1H), 8.00-7.40 (m, 2H), 7.86 (d, 1H), 7.58 (t, 1H), 7.35-7.27 (m, 2H), 4.56 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{16}\text{N}_5\text{O}_2\text{F}_3$: 416 (MH^+).
- [0426] 3-amino-6-[3-([(3-chloro-2,6-difluorophenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.05 (t, 1H), 8.89 (s, 1H), 8.83 (q, 1H), 8.48-8.46 (m, 1H), 8.37 (d, 1H), 7.90-7.20 (m, 2H), 7.83 (d, 1H), 7.66-7.58 (m, 1H), 7.55 (t, 1H), 7.20 (td, 1H), 4.59 (d, 2H), 2.84 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{16}\text{N}_5\text{O}_2\text{F}_2\text{Cl}$: 432 (MH^+).
- [0427] 3-amino-6-{3-[(9H-fluoren-9-ylamino)carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.18 (d, 1H), 8.87 (s, 1H), 8.83 (q, 1H), 8.57-8.55 (m, 1H), 8.44 (d, 1H), 8.10-7.20 (m, 2H), 8.00 (d, 1H), 7.92 (d, 2H), 7.62-7.56 (m, 3H), 7.46 (t, 2H), 7.38-7.32 (m, 2H), 6.34 (d, 1H), 2.83 (d, 3H); MS (EI) for $\text{C}_{26}\text{H}_{21}\text{N}_5\text{O}_2$: 436 (MH^+).
- [0428] 3-amino-N-methyl-6-[3-([(2-methylphenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.01 (t, 1H), 8.91 (s, 1H), 8.83 (q, 1H), 8.55-8.53 (m, 1H), 8.39 (d, 1H), 8.00-7.30 (m, 2H), 7.90 (d, 1H), 7.58 (t,

1H), 7.32-7.26 (m, 1H), 7.20-7.14 (m, 3H), 4.52 (d, 2H), 2.85 (d, 3H), 2.35 (s, 3H); MS (EI) for $C_{21}H_{21}N_5O_2$: 376 (MH^+).

[0429] 3-amino-6-[3-(((3,4-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.13 (t, 1H), 8.91 (s, 1H), 8.87 (q, 1H), 8.56-8.53 (m, 1H), 8.37 (dt, 1H), 8.20-7.30 (m, 2H), 7.88 (dt, 1H), 7.57 (t, 1H), 7.14-7.04 (m, 3H), 4.45 (d, 2H), 2.85 (d, 3H), 2.20 (s, 3H), 2.19 (s, 3H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390 (MH^+).

[0430] 3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.07 (t, 1H), 8.92 (s, 1H), 8.85 (q, 1H), 8.56-8.54 (m, 1H), 8.40 (d, 1H), 8.00-7.30 (m, 2H), 7.89 (dt, 1H), 7.59 (t, 1H), 7.22 (dd, 1H), 7.07 (dd, 1H), 7.00 (td, 1H), 4.49 (d, 2H), 2.85 (d, 3H), 2.32 (s, 3H); MS (EI) for $C_{21}H_{20}N_5O_2F$: 394 (MH^+).

[0431] 3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.28 (t, 1H), 8.92 (s, 1H), 8.89 (q, 1H), 8.58-8.55 (m, 1H), 8.39 (d, 1H), 8.00-7.30 (m, 2H), 7.88 (d, 1H), 7.58 (t, 1H), 7.46-7.36 (m, 2H), 7.24-7.19 (m, 1H), 4.51 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{17}N_5O_2F_2$: 398 (MH^+).

[0432] 3-amino-6-[3-(((2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.13 (t, 1H), 8.91 (s, 1H), 8.84 (q, 1H), 8.55-8.52 (m, 1H), 8.40 (d, 1H), 8.00-7.10 (m, 2H), 7.89 (d, 1H), 7.58 (t, 1H), 7.43 (td, 1H), 7.36-7.29 (m, 1H), 7.24-7.15 (m, 2H), 4.57 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{18}N_5O_2F$: 380 (MH^+).

[0433] 3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.03 (t, 1H), 8.86 (s, 1H), 8.80 (q, 1H), 8.45 (t, 1H), 8.35 (dt, 1H), 8.00-7.28 (m, 2H), 7.81 (dt, 1H), 7.53 (t, 1H), 7.50-7.39 (m, 1H), 7.17-7.09 (m, 1H), 4.58 (d, 2H), 2.84 (d, 3H); MS (EI) for $C_{20}H_{16}N_5O_2F_3$: 416 (MH^+).

[0434] 3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.17 (t, 1H), 8.89 (s, 1H), 8.83 (q, 1H), 8.51 (t, 1H), 8.37 (dt, 1H), 8.00-7.00 (m, 2H), 7.85 (dt, 1H), 7.56 (t, 1H), 7.33-7.25 (m, 2H), 4.50 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{20}H_{16}N_5O_2F_3$: 416 (MH^+).

- [0435] '3-amino-6-{3-[[[4-(aminosulfonyl)phenyl]methyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.24 (t, 1H), 8.92 (s, 1H), 8.84 (q, 1H), 8.56 (s, 1H), 8.40 (d, 1H), 7.88 (d, 1H), 7.80 (d, 2H), 7.58 (dd, 1H), 7.54 (d, 2H), 7.34 (s, 2H), 4.60 (d, 2H), 2.83 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{20}\text{N}_6\text{O}_4\text{S}$: 441 (MH^+).
- [0436] '3-amino-6-[3-((2,3-dimethylphenyl)methyl)amino]carbonyl}phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.64 (s, 1H), 8.36 (s, 1H), 7.99 (m, 2H), 7.68 (d, 1H), 7.51 (dd, 1H), 7.17 (m, 3H), 7.23 (t, 1H), 4.70 (d, 2H), 3.03 (d, 3H), 2.35 (s, 3H), 2.29 (s, 3H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390 (MH^+).
- [0437] '3-amino-6-{3-[[[4-hydroxy-3-(methyloxy)phenyl]methyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -methanol): 8.76 (s, 1H), 8.54 (s, 1H), 8.20 (d, 1H), 7.86 (d, 1H), 7.57 (dd, 1H), 6.98 (s, 1H), 6.84 (d, 1H), 6.76 (d, 1H), 4.54 (s, 2H), 3.85 (s, 3H), 2.97 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_4$: 408 (MH^+).
- [0438] '3-amino-6-[3-[[[4-chloro-2-methylphenyl]methyl]amino]carbonyl}phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -methanol): 8.76 (s, 1H), 8.55 (s, 1H), 8.22 (d, 1H), 7.86 (d, 1H), 7.58 (dd, 1H), 7.29 (d, 1H), 7.23 (s, 1H), 7.17 (d, 1H), 4.60 (s, 2H), 2.97 (s, 3H), 2.40 (s, 3H); MS(EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{O}_2\text{Cl}$: 410 (MH^+).
- [0439] '3-amino-6-{3-[[[2-(ethyloxy)phenyl]methyl]amino]carbonyl}phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d-chloroform): 8.64 (s, 1H), 8.36 (s, 1H), 7.99 (m, 2H), 7.68 (d, 1H), 7.51 (dd, 1H), 7.37 (d, 1H), 7.28 (dd, 1H), 6.94 (dd, 1H), 6.87 (d, 1H), 6.82 (t, 1H), 4.69 (d, 2H), 4.13 (q, 2H), 3.03 (d, 3H), 1.45 (t, 3H); MS(EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_3$: 406 (MH^+).
- [0440] 3-amino-N-methyl-6-(3-[[[naphthalen-2-ylmethyl]amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz; d_6 -DMSO): 9.17 (br s, 1H), 8.9 (s, 1H), 8.82 (s, 1H), 8.56 (s, 1H), 8.42-8.36 (d, 1H), 8.24-8.18 (d, 1H), 8-7.82 (td, 3H), 7.64-7.46 (m, 5H), 5.00 (s, 2H), 2.57 (s, 3H); MS (EI) for $\text{C}_{24}\text{H}_{21}\text{N}_5\text{O}_2$: 412 (MH^+).
- [0441] 3-amino-N-methyl-6-(3-[[[tetrahydrofuran-2-ylmethyl]amino]carbonyl}phenyl)pyrazine-2-carboxamide: MS(EI) for $\text{C}_{18}\text{H}_{21}\text{N}_5\text{O}_3$: 356 (MH^+).
- [0442] 3-amino-6-[3-((2-chloro-6-fluorophenyl)methyl)amino]carbonyl}phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.90 (s, 1H), 8.82 (s,

1H), 8.85 (s, 1H), 8.34-8.38 (d, 1H), 7.88-7.80 (d, 1H), 7.60-7.52 (t, 1H), 7.46-7.34 (m, 1H), 7.30-7.22 (t, 1H), 4.50 (s, 2H), 2.80 (s, 3H); MS (EI) for C₂₀H₁₇N₅O₂FCl : 414 (MH⁺).

[0443] 3-amino-6-{3-[[[3-fluoro-5-(trifluoromethyl)phenyl]methyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.28 (br s, 1H), 8.90 (s, 1H), 8.82 (s, 1H), 8.54 (s, 1H), 8.44-8.36 (d, 2H), 7.94-7.84 (d, 1H), 7.68-7.48 (m, 4H); MS (EI) for C₂₁H₁₇N₅O₂F₄: 448 (MH⁺).

[0444] 3-amino-6-{3-[[[2-[3,5-bis(methoxy)phenyl]ethyl]amino]carbonyl}phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.90 (s, 1H), 8.84 (d, 1H), 8.64 (s, 1H), 8.45 (s, 1H), 8.38-8.34 (d, 1H), 7.86-7.78 (d, 1H), 7.58-7.52 (t, 3H), 6.90-6.84 (d, 1H), 6.82-6.78 (d, 1H); MS (EI) for C₂₃H₂₅N₅O₄: 436 (MH⁺).

[0445] 3-amino-6-(3-[[[1,3-benzodioxol-5-ylmethyl]amino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.20 (s, 1H), 8.90 (s, 1H), 8.88-8.82 (d, 1H), 8.56 (s, 1H), 8.40-8.32 (d, 1H), 7.90-7.80 (d, 1H), 7.60-7.50 (t, 1H), 6.95 (s, 1H), 6.90-6.78 (q, 2H), 6.00 (s, 2H), 4.50-4.40 (d, 2H), 2.85 (s, 3H); MS (EI) for C₂₁H₁₉N₅O₄: 406 (MH⁺).

[0446] 3-amino-N-methyl-6-(3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.20 (t, 1H), 8.88 (s, 1H), 8.86-8.78 (d, 1H), 8.54 (s, 1H), 8.42-8.36 (d, 1H), 7.90-7.84 (d, 1H), 7.64-7.58 (t, 1H), 7.52-7.46 (d, 2H), 7.38-7.32 (d, 2H), 4.56-4.50 (d, 2H), 2.90-2.80 (t, 3H); MS (EI) for C₂₁H₁₈N₅O₃F₃: 446 (MH⁺).

[0447] 3-amino-6-{3-[[[4-fluoro-3-(trifluoromethyl)phenyl]methyl]aminocarbonyl}phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.24-9.18 (t, 1H), 8.90 (s, 1H), 8.86-8.80 (d, 1H), 8.52 (s, 1H), 8.42-8.37 (d, 1H), 7.90-7.84 (d, 1H), 7.80-7.72 (m, 2H), 7.62-7.58 (q, 1H), 7.54-7.48 (t, 1H), 4.60-4.55 (d, 2H), 3.78-3.68 (d, 3H); MS (EI) for C₂₁H₁₇N₅O₂F₄: 448 (MH⁺).

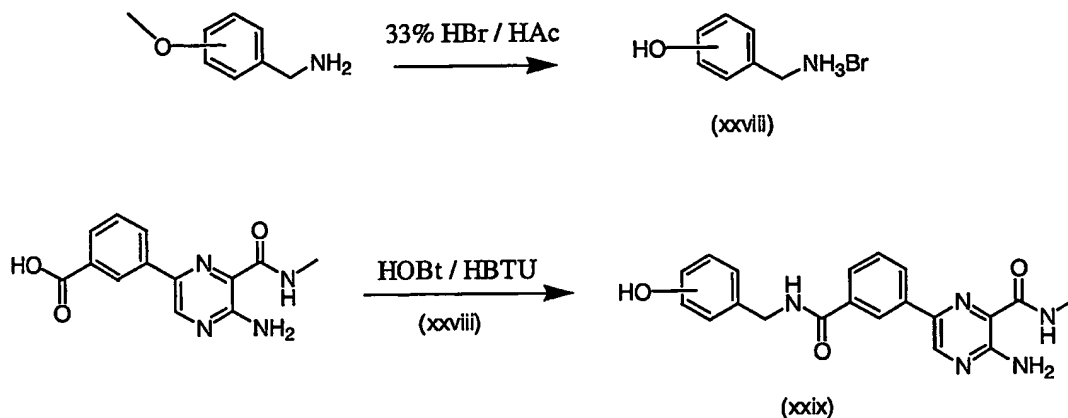
[0448] 3-amino-6-{3-[[[4-chloro-3-(trifluoromethyl)phenyl]methyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.28-9.22 (t, 1H), 8.90 (s, 1H), 8.88-8.82 (d, 1H), 8.52 (s, 1H), 8.44-8.36 (d, 1H), 7.90-7.82 (m, 2H), 7.74-7.66 (q, 2H), 7.60-7.54 (t, 1H); MS (EI) for C₂₁H₁₇N₅O₂F₃Cl: 464 (MH⁺).

Example 5

3-amino-6-[3-(hydroxy-benzylcarbamoyl)-phenyl]-pyrazine-2-carboxylic acid methylamide isomers

[0449] Scheme 5 shows how 3-amino-6-[3-(hydroxy-benzylcarbamoyl)-phenyl]-pyrazine-2-carboxylic acid methylamide isomers were made. Methoxy benzylamines were demethylated via acidic hydrolysis to give the corresponding amine hydrobromide salts, (xxviii). These were used to make compounds (xxix) via amide forming reaction, for example using standard amide coupling reagents as depicted in Scheme 5. Exemplary details are set forth below.

Scheme 5



[0450] 3-amino-6-(3-[(2-hydroxybenzyl)amino]carbonyl)phenyl)-N-methyl-2-pyrazinecarboxamide trifluoroacetate: A solution of 3.0 mL (0.023 mol) {2-(methoxy)phenyl}methyl}amine was refluxed in 50 mL 33% HBr in glacial acetic acid until the full deprotection of the starting material. The solvent was evaporated to obtain 4.7 g 2-(aminomethyl)phenol hydrobromide: MS (EI) for C₇H₉NO: 124.1 (MH⁺).

[0451] To a solution of 54 mg (0.20 mmol) 3-{5-amino-6-[(methylamino)carbonyl]-2-pyrazinyl}benzoic acid in DMF (5mL), 165 mg (1.20 mmol) 1-hydroxybenzotriazol (HOBt), 380 mg (1.00 mmol) O-benzotriazol-1-yl-N,N,N',N'-tetramethyluronium hexafluorophosphate (HBTU) and 0.22 mL (2.00 mmol) 4-methylmorpholine was added and the reaction mixture was stirred for 30 minutes at room temperature, followed by the addition of 41 mg (0.20 mmol) 2-(aminomethyl)phenol hydrobromide. The reaction mixture was stirred at room temperature for 18 h. The reaction was diluted with ethyl

acetate washed sequentially with water, 1.0 M aqueous hydrochloric acid, water and saturated aqueous sodium chloride. The organic layer was dried over sodium sulfate, filtered, and concentrated on a rotary evaporator under reduced pressure. The crude product was purified by HPLC (reverse-phase, CH₃CN/H₂O with 0.1% TFA). Upon removal of solvent, the product was taken up in methanol. Removal of solvent *in vacuo* provided 54 mg (55% yield) of title product: ¹H NMR (400 MHz, d₆-DMSO): 9.02 (t, 1H), 8.84 (s, 1H), 8.78 (q, 1H), 8.50 (s, 1H), 8.36 (d, 1H), 7.78 (d, 1H), 7.54 (t, 1H), 7.22 (m, 2H), 6.94 (m, 2H), 4.86 (bs, 1H), 4.40 (d, 2H), 2.80 (d, 3H). MS (EI) for C₂₀H₁₉N₅O₃: 378.5 (MH⁺).

[0452] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0453] 3-amino-6-(3-([(3-hydroxybenzyl)amino]carbonyl)phenyl)-N-methyl-2-pyrazine-carboxamide trifluoroacetate: ¹H NMR (400 MHz, d₆-DMSO): 9.02 (t, 1H), 8.86 (s, 1H), 8.80 (q, 1H), 8.54 (s, 1H), 8.36 (d, 1H), 7.86 (d, 1H), 7.58 (t, 1H), 7.15 (m, 2H), 6.94 (s, 1H), 6.76 (m, 1H), 4.94 (bs, 1H), 4.43 (d, 2H), 2.82 (d, 3H). MS (EI) for C₂₀H₁₉N₅O₃: 378.4 (MH⁺).

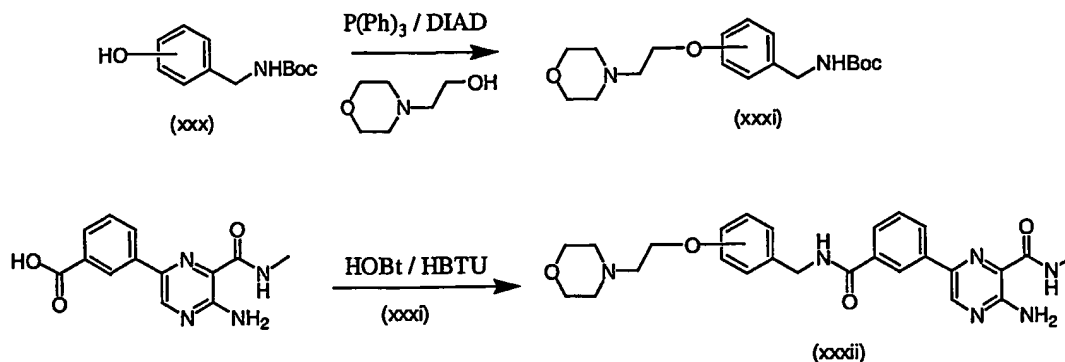
[0454] 3-amino-6-(3-([(4-hydroxybenzyl)amino]carbonyl)phenyl)-N-methyl-2-pyrazine-carboxamide trifluoroacetate: ¹H NMR (400 MHz, d₆-DMSO): 9.04 (t, 1H), 8.88 (s, 1H), 8.84 (q, 1H), 8.54 (s, 1H), 8.40 (d, 1H), 7.86 (d, 1H), 7.60 (t, 1H), 7.18 (d, 2H), 6.86 (d, 2H), 5.02 (bs, 1H), 4.43 (d, 2H), 2.82 (d, 3H). MS (EI) for C₂₀H₁₉N₅O₃: 378.4 (MH⁺).

Example 6

3-amino-N-methyl-6-{3-[(2-[2-(4-morpholinyl)ethoxy]benzyl)amino]carbonyl}phenyl}-2-pyrazinecarboxamide

[0455] Scheme 6 shows how 3-Amino-6-{3-[3-(morpholin-4-yl-ethoxy)-benzylcarbonyl]phenyl}-pyrazine-2-carboxylic acid methylamide isomers were made. Benzylamine hydrobromide salts (xxviii) (refer to Scheme 5) were converted to the corresponding Boc-protected derivatives (xxx). These were used to make compounds (xxxi), for example via Mitsunobu reaction. Intermediates (xxxi) were used to make compounds (xxxii) via amide forming reaction, for example using standard amide coupling reagents as depicted in Scheme 6. Exemplary details are set forth below.

Scheme 5



[0456] 3-amino-N-methyl-6-{3-[(2-[2-(4-morpholinyl)ethoxy]benzyl)amino)carbonyl]phe-nyl}-2-pyrazinecarboxamide: To an ice-cold solution of 4.0 g (0.02 mol) 2-(aminomethyl)phenol hydrobromide was added 8.40 g (0.10 mol) sodium bicarbonate in 10% aqueous THF (150mL) followed by 22 mL (0.022 mol) di-tert-butyl dicarbonate (1M in THF) and the reaction mixture was stirred at room temperature for 18 h. It was diluted with ethyl acetate and washed sequentially with water, 1.0 M aqueous hydrochloric acid, water and saturated aqueous sodium chloride. The organic layer was dried over sodium sulfate, filtered, and concentrated on a rotary evaporator under reduced pressure. The solvent was evaporated to obtain 4.2 g 1,1-dimethylethyl-[(2-hydroxyphenyl)-methyl]carbamate. MS (EI) for C₁₂H₁₇NO₃: 123.4 (M-Boc), 167.4 (M-tBu).

[0457] To an ice-cold solution of 3.30 g (0.015 mol) 1,1-dimethylethyl-[(2-hydroxyphenyl)-methyl]carbamate, 2.20 mL (0.018 mol) 2-morpholin-4-ylethanol and 4.80 g (0.019 mol) triphenylphosphine in a mixture of tetrahydrofuran-dichloromethane (2:1) (150 mL), 3.74 mL (0.019 mol) diisopropyl azodicarboxylate was added dropwise. The reaction mixture was stirred for 18h. It was diluted with ethyl acetate and washed sequentially with water then saturated aqueous sodium chloride. The organic layer was dried over sodium sulfate, filtered and concentrated on a rotary evaporator under reduced pressure. The resulting crude material was purified by silica gel column chromatography to yield 3.6 g 1,1-dimethylethyl-[(2-{[2-(4-morpholin-4-ylethyl)oxy]phenyl}methyl) carbamate, 71%. MS (EI) for C₁₈H₂₈N₂O₄: 337.4 (MH⁺), 236.4 (M-Boc).

[0458] Weight, 0.35g (1.0mmol), of 1,1-dimethylethyl-[(2-{[2-(4-morpholin-4-ylethyl)oxy]phenyl}methyl) carbamate was taken into dichloromethane (2mL) and treated with 2.0 mL trifluoroacetic acid for 30 minutes at room temperature. The solvent was

evaporated and the resulting crude ({2-[(2-morpholin-4-ylethyl)oxy]phenyl}methyl) trifluoroacetate, MS (EI) for $C_{13}H_{20}N_2O_2$: 337.4 (MH^+) was coupled with 0.27 g (1.00 mmol) 3-{5-amino-6-[(methylamino)carbonyl]-2-pyrazinyl}benzoic acid as described above. The reaction was diluted with ethyl acetate, washed sequentially with water, and saturated aqueous sodium chloride. The organic layer was dried over sodium sulfate, filtered and concentrated on a rotary evaporator under reduced pressure. The crude product was purified by HPLC (reverse-phase, CH_3CN/H_2O with 0.1% TFA) Upon removal of solvent, the product was taken up in a mixture of 10% methanol in ethyl acetate (100 mL) and washed with saturated aqueous sodium bicarbonate (2x 30 mL) then brine. The organic layer was dried over anhydrous sodium sulfate, filtered and concentrated *in vacuo* to provide 0.22 g title product, 44% yield. 1H NMR (400 MHz, d_6 -DMSO): 9.04 (t, 1H), 8.84 (s, 1H), 8.80 (q, 1H), 8.54 (s, 1H), 8.38 (d, 1H), 7.82 (d, 1H), 7.60 (t, 1H), 7.18 (m, 2H), 6.96 (m, 2H), 4.44 (d, 2H), 4.04 (m, 2H), 3.85 (t, 4H), 2.84 (d, 3H) 2.78 (m, 2H), 2.37 (t, 4H). MS (EI) for $C_{26}H_{30}N_6O_4$: 491.5 (MH^+).

[0459] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0460] 3-amino-N-methyl-6-{3-[(3-[2-(4-morpholinyl)ethoxy]benzyl)amino)carbonyl]phenyl}-2-pyrazinecarboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.03 (t, 1H), 8.86 (s, 1H), 8.84 (q, 1H), 8.54 (s, 1H), 8.38 (d, 1H), 7.84 (d, 1H), 7.58 (t, 1H), 7.14 (m, 2H), 6.92 (s, 1H), 6.74 (m, 1H), 4.40 (d, 2H), 4.02 (m, 2H), 3.84 (t, 4H), 2.80 (d, 3H) 2.76 (m, 2H), 2.34 (t, 4H). MS (EI) for $C_{26}H_{30}N_6O_4$: 491.5 (MH^+).

[0461] 3-amino-N-methyl-6-{3-[(4-[2-(4-morpholinyl)ethoxy]benzyl)amino)carbonyl]phenyl}-2-pyrazinecarboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.02 (t, 1H), 8.88 (s, 1H), 8.84 (q, 1H), 8.52 (s, 1H), 8.37 (d, 1H), 7.82 (d, 1H), 7.58 (t, 1H), 7.16 (d, 2H), 6.84 (d, 2H), 4.42 (d, 2H), 4.06 (m, 2H), 3.82 (t, 4H), 2.82 (d, 3H) 2.78 (m, 2H), 2.36 (t, 4H). MS (EI) for $C_{26}H_{30}N_6O_4$: 491.5 (MH^+).

[0462] 3-amino-6-(3-[(2-methoxybenzyl)amino]carbonyl)phenyl)-N-methyl-2-pyrazinecarboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.03 (t, 1H), 8.86 (s, 1H), 8.84 (q, 1H), 8.52 (s, 1H), 8.40 (d, 1H), 7.82 (d, 1H), 7.56 (t, 1H), 7.20 (m, 2H), 6.96 (m, 2H), 4.44 (d, 2H), 3.64 (s, 3H), 2.84 (d, 3H). MS (EI) for $C_{21}H_{21}N_5O_3$: 392.4 (MH^+).

[0463] 3-amino-N-methyl-6-[3-({[4-(4-methyl-1-piperazinyl)benzyl]amino}carbonyl)phenyl]-2-pyrazinecarboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.04 (t, 1H), 8.86 (s, 1H), 8.82 (dd, 1H), 8.52 (s, 1H), 8.38 (d, 1H), 7.84 (d, 1H), 7.58 (t, 1H), 7.20 (d, 2H), 6.84 (d, 2H), 4.43 (d, 2H), 3.08 (t, 4H), 2.82 (d, 3H), 2.42 (t, 4H), 2.20 (s, 3H). MS (EI) for $\text{C}_{25}\text{H}_{29}\text{N}_7\text{O}_2$: 460.2 (MH^+).

Example 7

3-amino-6-[3-({[4-(4-aminophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide

[0464] A solution of 3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid (0.10 g, 0.37 mmol), HOBt (0.055 g, 0.41 mmol) and HBTU (0.15 g, 0.41 mmol) in DMF (1.2 mL) was stirred at r.t. for 10 min. A solution of [4-(aminomethyl)phenyl]amine (0.050 mL, 0.44 mmol) in DMF (1.0 mL) was added and the resulting solution was stirred for 12h at r.t. The solution was concentrated *in vacuo* and the crude mixture was partitioned between EtOAc (100 mL) and brine (50 mL). The organic layer was washed with brine (2 x 50 mL) and the organic layer was dried (Na_2SO_4), filtered and concentrated *in vacuo*. The residue was purified via column chromatography (SiO_2 , 5% MeOH/ CH_2Cl_2) and trituration of the product gave 0.035 g (39% yield) of product: ^1H NMR (400 MHz, d_6 -DMSO): 8.95 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.50 (m, 1H), 8.37 (dt, 1H), 7.87 (dt, 1H), 7.56 (t, 1H), 7.01 (d, 2H), 6.52 (m, 2H), 4.97 (s, 2H), 4.35 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{20}\text{N}_6\text{O}_2$: 377 (MH^+).

[0465] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0466] 3-amino-6-[3-({[(2-aminophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide. ^1H NMR (400 MHz, d_6 -DMSO): 9.02 (t, 1H), 8.90 (s, 1H), 8.84 (m, 1H), 8.51 (m, 1H), 8.38 (dt, 1H), 7.88 (dt, 1H), 7.58 (t, 1H), 7.07 (m, 1H), 6.97 (dt, 1H), 6.63 (dd, 1H), 6.52 (dt, 1H), 5.19 (s, 2H), 4.38 (d, 2H), 2.85 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{20}\text{N}_6\text{O}_2$: 377 (MH^+).

[0467] 3-amino-6-[3-({[(3-aminophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide. ^1H NMR (400 MHz, d_6 -DMSO): 9.03 (t, 1H), 8.91 (s, 1H), 8.84 (m, 1H), 8.53 (m, 1H), 8.39 (dt, 1H), 7.89 (dt, 1H), 7.58 (t, 1H), 6.96 (t, 1H),

6.54 (m, 1H), 6.48 (d, 1H), 6.43 (m, 1H), 5.05 (s, 2H), 4.40 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{20}H_{20}N_6O_2$: 377 (MH^+).

[0468] 3-amino-6-[3-([4-(dimethylamino)phenyl]methyl)amino]carbonylphenyl]-*N*-methylpyrazine-2-carboxamide. 1H NMR (400 MHz, d_6 -DMSO): 9.01 (t, 1H), 8.91 (s, 1H), 8.85 (m, 1H), 8.51 (s, 1H), 8.37 (d, 1H), 7.87 (d, 1H), 7.56 (t, 1H), 7.18 (d, 2H), 6.70 (d, 2H), 4.41 (d, 2H), 2.85 (m, 9H); MS (EI) for $C_{22}H_{24}N_6O_2$: 405 (MH^+).

[0469] 3-amino-6-[3-([(3-bromophenyl)methyl]amino)carbonylphenyl]-*N*-methylpyrazine-2-carboxamide. 1H NMR (400 MHz, d_6 -DMSO): 9.19 (t, 1H), 8.91 (s, 1H), 8.85 (m, 1H), 8.53 (m, 1H), 8.39 (dt, 1H), 7.89 (dt, 1H), 7.59 (t, 1H), 7.56 (s, 1H), 7.47 (m, 1H), 7.37 (d, 1H), 7.32 (t, 1H), 4.53 (d, 2H), 2.86 (d, 3H); MS (EI) for $C_{20}H_{18}N_5O_2Br$: 440 (M^+).

[0470] 3-amino-6-[3-([(4-bromophenyl)methyl]amino)carbonylphenyl]-*N*-methylpyrazine-2-carboxamide. 1H NMR (400 MHz, d_6 -DMSO): 9.19 (t, 1H), 8.91 (m, 1H), 8.85 (m, 1H), 8.53 (m, 1H), 8.39 (m, 1H), 7.88 (m, 1H), 7.58 (t, 1H), 7.54 (d, 2H), 7.34 (d, 2H), 4.50 (d, 2H), 2.86 (d, 3H).

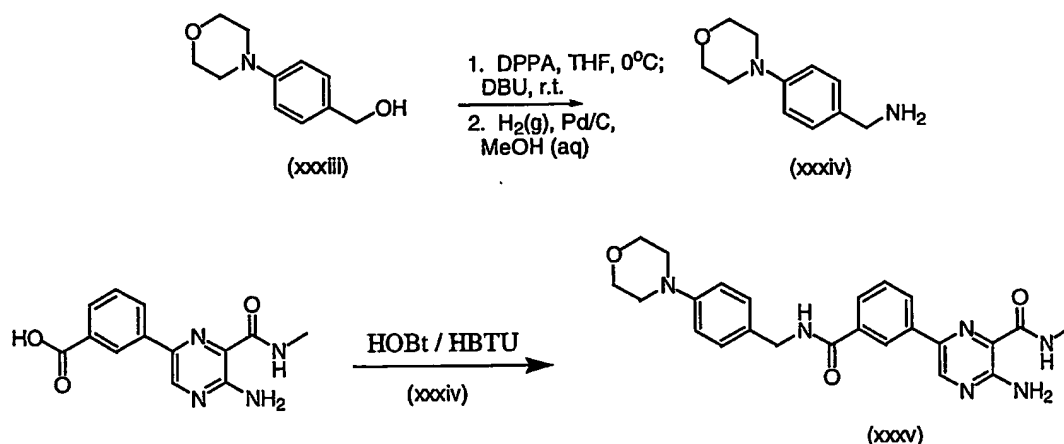
[0471] 3-amino-6-[3-([(2-bromophenyl)methyl]amino)carbonylphenyl]-*N*-methylpyrazine-2-carboxamide. 1H NMR (400 MHz, d_6 -DMSO): 9.20 (m, 1H), 8.93 (m, 1H), 8.86 (m, 1H), 8.58 (m, 1H), 8.41 (d, 1H), 7.91 (m, 1H), 7.64 (d, 1H), 7.60 (t, 1H), 7.39 (m, 2H), 7.24 (m, 1H), 4.57 (d, 2H), 2.85 (d, 3H); MS (EI) for $C_{20}H_{18}N_5O_2Br$: 442 ($M^+ + 2$).

Example 8

3-amino-N-methyl-6-[3-([(3-morpholin-4-ylphenyl)methyl]amino)methyl]phenyl]pyrazine-2-carboxamide

[0472] Scheme 7 shows how 3-amino-N-methyl-6-[3-([(3-morpholin-4-ylphenyl)methyl]amino)methyl]phenyl pyrazine-2-carboxamide was made. (4-Morpholin-4-yl-phenyl)-methanol (xxxiii) was converted to the corresponding amine (xxxiv). Amine (xxxiv) was used to make compound (xxxv) via amide forming reaction, for example using standard amide coupling reagents as depicted in Scheme 7. Exemplary details and further analogous examples are set forth below.

Scheme 7



[0473] A solution of (3-morpholin-4-ylphenyl)methanol (0.10 g, 0.52 mmol), in THF (1.0 mL) was cooled to 0°C and DPPA (0.13 mL, 0.62 mmol) was added, followed by DBU (0.093 mL, 0.62 mmol). The solution was allowed to warm to r.t. and stirred for 12h. The solution was poured into EtOAc (100 mL) and this was washed with H₂O (50 mL) and 5% HCl(aq) (50 mL). The organic layer was dried (Na₂SO₄), filtered and concentrated *in vacuo* to give 0.16 g (>100% yield) of 4-[3-(azidomethyl)phenyl]morpholine, (TLC: UV active spot, R_f of 0.7 in 1:1 hexanes/EtOAc), which was used without further purification.

[0474] To a solution of 10% Pd/Carbon (0.030 g, 0.029 mmol) in 10% aqueous MeOH (4.2 mL) was added 4-[3-(azidomethyl)phenyl]morpholine (0.16 g, 0.71 mmol) in MeOH (4.2 mL), and the solution was stirred vigorously under a H₂(g) balloon for 3 h. The solution was filtered through celite/fritted glass funnel and concentrated *in vacuo*. The crude residue was purified via HPLC (reverse-phase, CH₃CN/H₂O with 0.1% TFA) and fractions containing the desired product were concentrated *in vacuo*, to provide 0.064 g (29% yield) of [(3-morpholin-4-ylphenyl)methyl]amine trifluoroacetate.

[0475] A solution of 3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoic acid (0.081 g, 0.30 mmol), HOBt (0.048 g, 0.36 mmol) and HBTU (0.17 g, 0.45 mmol) in DMF (1.0 mL) was stirred at r.t. for 10 min. A solution of [(3-morpholin-4-ylphenyl)methyl]amine trifluoroacetate (0.13 g, 0.30 mmol) in DMF (1.0 mL) was added

and the resulting solution was stirred for 12 h at r.t. The solution was concentrated *in vacuo* and the crude mixture was purified via HPLC (reverse-phase, CH₃CN/H₂O with 0.1% TFA). Upon removal of CH₃CN/H₂O, the product was taken up in EtOAc (100 mL) and washed with 1M NaOH(aq) (50 mL). The organic layer was dried (Na₂SO₄), filtered and concentrated *in vacuo*. Trituration and collection of a solid by vacuum filtration provided 11.6 mg (and treated with Bio-Rad AG 1-X8 resin (hydroxide form) until pH 8. The product was filtered and concentrated *in vacuo*, to provide 14.1 mg (9% yield) of 3-amino-*N*-methyl-6-[3-({[(3-morpholin-4-ylphenyl)methyl]amino}methyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.15 (t, 1H), 8.91 (s, 1H), 8.87 (m, 1H), 8.55 (s, 1H), 8.37 (d, 1H), 7.88 (d, 1H), 7.57 (t, 1H), 7.19 (t, 1H), 6.95 (s, 1H), 6.82 (m, 2H), 4.48 (d, 2H), 3.72 (t, 4H), 3.08 (t, 2H), 2.85 (d, 3H); MS (EI) for C₂₄H₂₆N₆O₃: 447 (MH⁺).

[0476] Using the analogous synthetic techniques beginning from commercially available [(2-morpholin-4-ylphenyl)methyl]amine and [(4-morpholin-4-ylphenyl)methyl]amine, the following compounds of the invention were prepared:

[0477] 3-amino-*N*-methyl-6-[3-({[(2-morpholin-4-ylphenyl)methyl]amino}methyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.06 (t, 1H), 8.92 (s, 1H), 8.85 (m, 1H), 8.56 (s, 1H), 8.39 (d, 1H), 7.91 (d, 1H), 7.58 (t, 1H), 7.31 (d, 1H), 7.25 (t, 1H), 7.15 (d, 1H), 7.09 (t, 1H), 4.65 (d, 2H), 3.77 (m, 4H), 2.89 (m, 2H), 2.85 (d, 3H); MS (EI) for C₂₄H₂₆N₆O₃: 447 (MH⁺).

[0478] 3-amino-*N*-methyl-6-[3-({[(4-morpholin-4-ylphenyl)methyl]amino}methyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.16 (t, 1H), 8.91 (s, 1H), 8.88 (m, 1H), 8.56 (s, 1H), 8.37 (dt, 1H), 7.88 (dt, 1H), 7.56 (t, 1H), 7.23 (d, 2H), 6.91 (d, 2H), 4.43 (d, 2H), 3.72 (m, 4H), 3.06 (m, 2H), 2.85 (d, 3H); MS (EI) for C₂₄H₂₆N₆O₃: 447 (MH⁺).

[0479] Methyl 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoate: A mixture of 3-amino-6-bromopyrazine-2-cyclopropylcarboxamide (5.0 g, 19.4 mmole), [1,1'-bis-(diphenylphosphino)ferrocene]dichloropalladium(II)-dichloromethane complex (0.88 g, 1.9 mmol), 3-carbomethoxyphenylboronic acid (4.2 g, 23.3 mmol) in DMF (30 mL) was de-gassed with nitrogen. Triethylamine (5.4 mL, 38.9 mmol) was then added and the resulting mixture was heated at 85 ° for 2.5 hours. The reaction mixture was cooled to room temperature, diluted with ethyl acetate (200 mL), filtered over silica gel, washed with saturated sodium bicarbonate solution (150 mL), dried over sodium sulfate,

filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give methyl 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoate (a tan color solid, 4.5 g, 74% yield): ^1H NMR (400 MHz, DMSO- d_6): 8.67 (s, 1H), 8.50-8.49 (t, 1H), 8.07-8.03 (m, 2H), 7.57-7.54 (t, 1H), 3.98 (s, 3H), 2.91-2.88 (m, 1H), 0.94-0.85 (m, 2H), 0.72-0.62 (m, 2H); MS (EI) for $\text{C}_{16}\text{H}_{16}\text{N}_4\text{O}_3$: 428 (MH^+).

[0480] Methyl 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoate: A mixture of 3-amino-6-bromopyrazine-2-ethylcarboxamide (5.0 g, 20.4 mmol), [1,1'-bis-(diphenylphosphino)ferrocene]dichloropalladium(II)-dichloromethane complex (0.93 g, 2.04 mmol), 3-carbomethoxyphenylboronic acid (4.4 g, 24.4 mmol) in DMF (30 mL) was de-gassed with nitrogen. Triethylamine (5.7 mL, 40.1 mmol) was then added and the resulting mixture was heated at 85 ° for 2.5 hours. The reaction mixture was cooled to room temperature, diluted with ethyl acetate (200mL), filtered over silica gel, washed with saturated sodium bicarbonate solution (150 mL), dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give methyl 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoate (4.3 g, 70% yield): ^1H NMR (400 MHz, CDCl_3): 8.66 (s, 1H), 8.53-8.52 (t, 1H), 8.08-8.06 (m, 2H), 8.00 (b, 1H), 3.5 (t, 1H), 3.97 (s, 3H), 2.91-2.88 (m, 1H), 0.94-0.85 (m, 2H), 0.72-0.62 (m, 2H); MS (EI) for $\text{C}_{15}\text{H}_{16}\text{N}_4\text{O}_3$: 301 (MH^+).

[0481] 3-amino-N-ethyl-6-(3-[(phenylmethyl)amino]carbonyl)phenylpyrazine-2-carboxamide: Methyl 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoate (480 mg, 1.6 mmol) was suspended in a 20 mL of 1:1 MeOH-5% aqueous sodium hydroxide and heated at 90-95°C for 4 hours. The homogeneous reaction mixture was cooled to room temperature, concentrated to half the initial volume, cooled in an ice-bath, and acidified to pH of 4 with concentrated HCl. A white solid precipitated and was collected, washed with water and dried *in vacuo* to give 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoic acid (445 mg, 97% yield). The pyrazine acid (40 mg, 0.14 mmol) was dissolved in DMF (1mL), and then 1-hydroxybenzotriazole (30 mg, 0.22 mmol), 1-[(3-dimethylamino)propyl]-3-ethylcarbodiimide hydrochloride (30 mg, 0.16 mmol), and 4-methylmorpholine (30 mg, 0.28 mol) were added sequentially. After one hour benzylamine (30mg, 0.27 mmol) was added to the reaction mixture and stirring was continued at room temperature for 18 hours. The mixture was

then diluted with ethyl acetate (10 mL), washed with water, saturated sodium bicarbonate solution then brine and dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give 3-amino-N-ethyl-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide (a tan color solid, 22 mg, 42% yield): ^1H NMR (400 MHz, DMSO- d_6): 9.14 (s, 1H), 8.88 (s, 1H), 8.87-8.85 (b, 1H), 8.51 (s, 1H), 8.37-8.34 (d, 1H), 7.87-7.85 (d, 1H), 7.63-7.59 (b, 1H), 7.58-7.54 (t, 1H), 7.34-7.23 (m, 4H), 4.53-4.52 (d, 2H), 3.38-3.29 (m, 2H), 1.17-1.14 (t, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376 (MH^+).

[0482] 3-amino-6-[3-((2-chloro-6-fluorophenyl)methyl)amino]carbonylphenyl]-N-ethylpyrazine-2-carboxamide: Methyl 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoate (480 mg, 1.6 mmol) was suspended in a 20 mL of 1:1 MeOH-5% aqueous sodium hydroxide and heated at 90-95°C for 4 hours. The homogeneous reaction mixture was cooled to room temperature, concentrated to half the initial volume, cooled in an ice-bath, and acidified to pH 4 with concentrated HCl. A white solid precipitated and was collected by filtration, washed with water and dried *in vacuo* to give 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoic acid (445 mg, 97% yield). The acid (40 mg, 0.14 mmol) was dissolved in DMF (1mL) and 1-hydroxybenzotriazole (30 mg, 0.22 mmol), 1-[(3-dimethylamino)propyl]-3-ethylcarbodiimide hydrochloride (30 mg, 0.16 mmol), and 4-methylmorpholine (30 mg, 0.28 mol) were added sequentially. After one hour, 2-chloro-6-fluorobenzylamine (30mg, 0.19 mmol) was added to the reaction mixture and stirring was continued at room temperature for 18 hours. It was then diluted with ethyl acetate (10 mL), washed with water, saturated sodium bicarbonate solution, and brine, dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give 3-amino-6-[3-((2-chloro-6-fluorophenyl)methyl)amino]carbonylphenyl]-N-ethylpyrazine-2-carboxamide (a tan color solid, 31 mg, 51% yield): ^1H NMR (400 MHz, DMSO- d_6): 8.86 (s, 1H), 8.85-8.83 (b, 1H), 8.45 (s, 1H), 8.33-8.31 (d, 1H), 7.83-7.81 (d, 1H), 7.53-7.51 (t, 1H), 7.39-7.34 (m, 2H), 7.26-7.21 (t, 1H), 4.63-4.61 (d, 2H), 3.38-3.29 (m, 2H), 1.15-1.11 (t, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{ClFN}_5\text{O}_2$: 428 (MH^+).

[0483] 3-amino-N-cyclopropyl-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide: Methyl 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoate

(480 mg, 1.5 mmol) was suspended in a 20 mL of 1:1 MeOH-5% aqueous sodium hydroxide and heated at 90-95 °C for 4 hours. The homogeneous reaction mixture was cooled to room temperature, concentrated to half the initial volume, cooled in an ice-bath, and acidified to pH 4 with concentrated HCl. A white solid precipitated and was collected by filtration, washed with water and dried *in vacuo* to give the intermediate 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoic acid. The acid (40 mg, 0.13 mmol) was dissolved in DMF (1 mL) and 1-hydroxybenzotriazole (30 mg, 0.22 mmol), 1-[(3-dimethylamino)propyl]-3-ethylcarbodiimide hydrochloride (30 mg, 0.16 mmol), and 4-methylmorpholine (30 mg, 0.28 mol) were added sequentially. After one hour benzylamine (30 mg, 0.27 mmol) was added to the reaction mixture and stirring was continued at room temperature for 18 hours. It was then diluted with ethyl acetate (10 mL), washed with water, saturated sodium bicarbonate solution, and brine, dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give 3-amino-N-cyclopropyl-6-(3-[(phenylmethyl)amino]carbonyl)phenyl pyrazine-2-carboxamide (a tan color solid, 23 mg, 44% yield): ¹H NMR (400 MHz, DMSO-d₆): 9.14 (t, 1H), 8.88 (s, 1H), 8.69-8.68 (d, 1H), 8.51 (s, 1H), 8.34-8.32 (d, 1H), 7.87-7.85 (d, 1H), 7.63-7.59 (b, 1H), 7.57-7.53 (t, 1H), 7.34-7.23 (m, 4H), 4.53-4.52 (d, 2H), 2.84-2.82 (m, 1H), 0.75-0.70 (m, 4H); MS (EI) for C₂₂H₂₁N₅O₂: 388 (MH⁺).

[0484] 3-amino-6-[3-([(2-chloro-6-fluorophenyl)methyl]amino)carbonyl]phenyl]-N-cyclopropylpyrazine-2-carboxamide: Methyl 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl} benzoate (480 mg, 1.5 mmol) was suspended in a 20 mL of 1:1 MeOH-5% aqueous sodium hydroxide and heated at 90-95°C for 4 hours. The homogeneous reaction mixture was cooled to room temperature, concentrated to half the initial volume, cooled in an ice-bath, and acidified to pH 4 with concentrated HCl. A white solid precipitated and was collected by filtration, washed with water and dried *in vacuo* to give the intermediate 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoic acid. The acid (40 mg, 0.13 mmol) was dissolved in DMF (1mL) and 1-hydroxybenzotriazole (30 mg, 0.22 mmol), 1-[(3-dimethylamino)propyl]-3-ethylcarbodiimide hydrochloride (30 mg, 0.16 mmol), and 4-methylmorpholine (30 mg, 0.28 mol) were added sequentially. After one hour 2-chloro-6-fluorobenzylamine (30mg, 0.19 mmol) was added to the reaction mixture and stirring was continued at room temperature for 18 hours. It was then diluted with ethyl acetate (10 mL), washed with water, saturated sodium bicarbonate solution, and brine, dried over sodium sulfate,

filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give 3-amino-6-[3-({[(2-chloro-6-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-cyclopropylpyrazine-2-carboxamide (a tan color solid, 23 mg, 56% yield): ^1H NMR (400 MHz, DMSO- d_6): 8.87-8.86 (t, 1H), 8.85-8.83 (b, 1H), 8.65-8.64 (d, 1H), 8.45 (s, 1H), 7.83-7.81 (d, 1H), 7.53-7.51 (t, 1H), 7.41-7.39 (m, 2H), 7.26-7.21 (t, 1H), 4.63-4.61 (d, 2H), 2.85-2.81 (m, 1H), 0.73-0.64 (m, 4H); MS (EI) for $\text{C}_{22}\text{H}_{19}\text{ClFN}_5\text{O}_2$: 440 (MH^+).

[0485] 3-amino-N-methyl-6-[3-[(2-phenylhydrazino)carbonyl]phenyl]pyrazine-2-carboxamide: 3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoic acid (40 mg, 0.15 mmol) was dissolved in DMF (1 mL) and 1-hydroxybenzotriazole (30 mg, 0.22 mmol), 1-[(3-dimethylamino)propyl]-3-ethylcarbodiimide hydrochloride (30 mg, 0.16 mmol), and 4-methylmorpholine (30 mg, 0.28 mol) were added sequentially. After one hour phenylhydrazine (30 mg, 0.27 mmol) was added to the reaction mixture and stirring was continued at room temperature for 18 hours. It was then diluted with ethyl acetate (10 mL), washed with water, saturated sodium bicarbonate solution, and brine, dried over sodium sulfate, filtered, and concentrated. The oily residue was purified by flash chromatography (silica gel, eluted with ethyl acetate-hexanes gradient from 15% to 80% yield) to give 3-amino-N-methyl-6-[3-[(2-phenylhydrazino)carbonyl]phenyl]pyrazine-2-carboxamide (a tan color solid, 30 mg, 56% yield): ^1H NMR (400 MHz, DMSO- d_6): 10.42 (b, 1H), 8.92 (s, 1H), 8.84-8.82 (b, 1H), 8.57-8.57 (s, 1H), 8.41-8.39 (d, 1H), 7.94-7.89 (m, 1H), 7.89-7.86 (d, 1H), 7.61-7.57 (t, 1H), 7.17-7.13 (t, 1H), 6.83-6.80 (d, 1H), 6.73-6.69 (m, 1H), 2.85 (s, 3H); MS (EI) for $\text{C}_{19}\text{H}_{18}\text{N}_6\text{O}_2$: 440 (MH^+).

Example 9

4-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoic acid

[0486] Phenylmethyl 4-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoate: To a stirred mixture of 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide (0.577g, 2.5 mmol), 4-benzyloxycarbonylphenylboronic acid (0.768 g, 3.0mmol), triethylamine (1.01 g, 10 mmol) in 10 mL of DMF was added [1,1'-bis(diphenylphosphino)-ferrocene]dichloropalladium(II) complex with dichloromethane (1:1), (20.4mg, 0.025 mmol). The stirred mixture was warmed to 80-90 °C for 12 h and then concentrated on a rotary evaporator under reduced pressure. The residue was dissolved in 50 mL of EtOAc,

washed with brine (30 mL). The organic layer was filtered through celite, and then dried over anhydrous sodium sulfate. Filtration, concentration and column chromatography on silica (EtOAc) gave a solid (450mg, 59.7% yield). ^1H NMR (400 MHz; DMSO- d_6): 8.91 (s, 1H); 8.84 (br s, 1H); 8.33 (d, 2H); 8.02 (d, 2H); 7.30-7.48 (m, 5H); 5.38 (s, 2H); 2.86 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_4\text{O}_3$: 363 (MH^+).

[0487] 4-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid: A flask was charged with phenylmethyl 4-[5-amino-6-(methylamino)carbonyl]pyrazin-2-yl}benzoate (0.37 g, 1.01 mmol) and 10 mL of a solution of 1 N aqueous LiOH. The mixture was refluxed for 15 min. and then allowed to cool to room temperature. The mixture was extracted with ethyl acetate (10mL). The pH of the aqueous phase was adjusted to pH of 2 – 3 with 10% aqueous HCl. The resulting solid product was collected by filtration, washed with water, and dried, to afford 4-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoic acid (0.150mg, 54.2% yield): ^1H NMR (400 MHz; DMSO- d_6): 8.93 (s, 1H); 8.80 (br s, 1H); 8.30 (d, 2H); 8.41 (d, 1H); 8.0 (d, 2H); 2.86 (d, 3H); MS (EI) for $\text{C}_{13}\text{H}_{12}\text{N}_4\text{O}_3$: 273 (MH^+).

[0488] 3-amino-N-methyl-6-(4-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- d_6): 9.18 (m, 1H); 8.95 (s, 1H); 8.90 (m, 1H); 8.3 (d, 2H); 8.0 (d, 2H); 7.37 (m, 5H); 4.5 (d, 2H); 2.87 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_5\text{O}_2$: 362 (MH^+).

[0489] 3-amino-N-methyl-6-(4-{[(2-phenylethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376 (MH^+)

[0490] 3-amino-6-[4-{[(2,4-dichlorophenyl)methyl]amino}carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- d_6): 9.18 (m, 1H); 8.97 (s, 1H); 8.90 (m, 1H); 8.3 (d, 2H); 8.0 (d, 2H); 7.62 (c, 1H); 7.4 (m, 2H); 4.53 (d, 2H); 2.87 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{Cl}_2$: 430 (MH^+)

[0491] 3-amino-N-methyl-6-[4-{[(4-methylphenyl)methyl]amino}carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- d_6): 9.1 (m, 1H); 8.97 (s, 1H); 8.90 (m, 1H); 8.3 (d, 2H); 8.0 (d, 2H); 7.2-7.1 (m, 5H); 4.53 (d, 2H); 2.87 (d, 3H); 2.25 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376 (MH^+)

[0492] 3-amino-6-(4-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_5\text{O}_2$: 388 (MH^+)

- [0493] 3-amino-N-methyl-6-[3-({[(2,4,6-trichlorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.86 (s, 1H); 8.8 (m, 1H); 8.68 (m, 1H); 8.42 (s, 1H); 8.34 (d, 1H); 7.8 (d, 1H); 7.54 (t, 1H); 7.46 (s, 1H); 7.34 (s, 1H); 4.6 (d, 2H); 2.85 (d, 3H); 2.43 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{16}\text{N}_5\text{O}_2\text{Cl}_3$: 465 (MH^+)
- [0494] 3-amino-6-[3-({[(3-chloro-2-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 9.25 (t, 1H); 8.88 (s, 1H); 8.82 (m, 1H); 8.5 (m, 1H); 8.38 (d, 1H); 7.86 (d, 1H); 7.56 (t, 1H); 7.34 (d, 1H); 7.28 (d, 1H); 7.2 (t, 1H); 4.5 (d, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{O}_2\text{Cl}$: 410 (MH^+)
- [0495] 3-amino-6-[3-({[(2,5-bis(methoxy)phenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.95 (t, 1H); 8.9 (s, 1H); 8.68 (m, 1H); 8.8 (m, 1H); 8.52 (s, 1H); 8.36 (d, 1H); 7.84 (d, 1H); 7.54 (t, 1H); 6.9 (d, 1H); 6.8 (m, 3H); 4.6 (d, 2H); 3.8 (s, 3H); 3.62 (s, 3H); 2.85 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 422 (MH^+)
- [0496] 3-amino-6-[3-({[(3-fluoro-4-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 9.1 (t, 1H); 8.85 (s, 1H); 8.8 (m, 1H); 8.52 (s, 1H); 8.38 (d, 1H); 7.85 (d, 1H); 7.58 (t, 1H); 7.2 (t, 1H); 4.45 (d, 2H); 2.85 (d, 3H); 2.2 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 394 (MH^+)
- [0497] 3-amino-6-[3-({[(2-chloro-6-fluoro-3-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.85 (s, 1H); 8.8 (m, 1H); 8.53 (s, 1H); 8.38 (d, 1H); 7.8 (d, 1H); 7.56 (t, 1H); 7.2 (m, 1H); 7.15 (t, 1H); 4.6 (d, 2H); 2.85 (d, 3H); 2.3 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_2\text{FCl}$: 428 (MH^+)
- [0498] 3-amino-6-[3-({[(2-chloro-3,6-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.9 (t, 1H); 8.85 (s, 1H); 8.8 (m, 1H); 8.49 (s, 1H); 8.38 (d, 1H); 7.8 (d, 1H); 7.56 (t, 1H); 7.45 (m, 1H); 7.3 (m, 1H); 4.6 (d, 2H); 2.85 (d, 3H); 2.3 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{16}\text{N}_5\text{O}_2\text{F}_2\text{Cl}$: 432 (MH^+)
- [0499] 3-amino-6-[3-({[(2,3-difluoro-4-methylphenyl)methyl]amino}carbonyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 9.1 (t, 1H); 8.9 (s, 1H); 8.8 (m, 1H); 8.53 (s, 1H); 8.38 (d, 1H); 7.8 (d, 1H); 7.56 (t, 1H); 7.1 (m, 1H); 4.5 (d, 2H); 2.85 (d, 3H); 2.23 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_2\text{F}_2$: 412 (MH^+)

Example 10

3-amino-N-methyl-6-{3-[(methylsulfonyl)amino]phenyl}pyrazine-2-carboxamide

[0500] 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide: To a solution of 3-amino-6-bromo-pyrazine-2-carboxylic acid methylamide (1.234 g, 5.30 mmol, 1.00 equiv.) in N,N-dimethylformamide (DMF, 30 mL) was added [1,1'-bis(diphenylphosphino)-ferrocene]dichloropalladium(II) complex with dichloromethane (1:1), (Aldrich Chemical Company, 433 mg, 0.530 mmol, 0.100 equiv.), 3-aminophenylboronic acid monohydrate (Aldrich Chemical Company, 1.149 g, 7.41 mmol, 1.40 equiv.), and triethylamine (TEA, 1.073 g, 10.6 mmol, 2.00 equiv.). Nitrogen was passed through the solution for 3-5 min. The reaction mixture was heated at 85-90 °C for 15 h with stirring. The reaction was allowed to cool to room temperature and diluted with 400 mL of EtOAc. The mixture was washed with saturated aqueous NaHCO₃ (2 X 50.0 mL), saturated aqueous NaCl (1 X 50.0 mL), dried over magnesium sulfate (MgSO₄), and filtered. The filtrate was concentrated on a rotary evaporator under reduced pressure. To the crude product was added 30 mL of a 4.0M solution of anhydrous HCl in dioxane. The resulting solid was filtered and rinsed with 20% methanol in diethyl ether to remove impurities. The solid was dissolved in methanol and neutralized with sufficient AG 1-X8 Resin (hydroxide form) that the pH of the solution became basic. The mixture was filtered, and the filtrate was concentrated on a rotary evaporator at reduced pressure to afford the title compound (0.695 g, 54% yield).

Example 11

3-amino-N-methyl-6-{3-[(methylsulfonyl)amino]phenyl}pyrazine-2-carboxamide

[0501] 3-amino-N-methyl-6-{3-[(methylsulfonyl)amino]phenyl}pyrazine-2-carboxamide: To a solution of 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide (26 mg, 0.11 mmol) in dry dichloromethane (2 mL) at 0° C were added triethylamine (22 mg, 0.22 mmol) and methanesulfonyl chloride (20 mg, 0.17 mmol). The reaction was slowly warmed to room temperature and stirred overnight. Reaction was treated water (10 mL) and diluted with ethyl acetate (100 mL). The organic layer was washed with saturated aqueous sodium bicarbonate (40 mL) and saturated aqueous sodium chloride (30 mL), dried over magnesium sulfate, filtered, and concentrated to afford 3-amino-N-methyl-6-{3-[(methylsulfonyl)amino]phenyl}pyrazine-2-carboxamide (15 mg, 45% yield) as

yellow solid. ^1H NMR (400 MHz, CDCl_3): 8.62 (s, 1H), 7.94 (br s, 1H), 7.75 (br s, 1H), 7.68 (br d, 1H), 7.45 (t, 1H), 7.30 (br d, 1H), 7.25 (br s, 1H), 3.02 (m, 6H); MS (EI) for $\text{C}_{13}\text{H}_{15}\text{N}_5\text{O}_3\text{S}$: 322 (MH^+).

Example 12

3-Amino-N-methyl-6-[3-({[4-(methyloxy)phenyl]carbonyl}amino)phenyl]pyrazine-2-carboxamide

[0502] To a solution of 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide (20 mg, 0.07 mmol) and triethylamine (160 mg, 0.16 mmol) in tetrahydrofuran (1 mL) was added p-anisoyl chloride (20 mg, 0.12 mmol) slowly at room temperature. The mixture was allowed to stir overnight. The reaction was quenched with water (10 mL) at 0° C and diluted with ethyl acetate (200 mL). The organic layer was washed with saturated aqueous sodium bicarbonate (30 mL), saturated aqueous sodium chloride, and dried over magnesium sulfate. The solution was filtered and concentrated at reduced pressure to afford 3-amino-N-methyl-6-[3-({[4-(methyloxy)phenyl]carbonyl}amino)phenyl]pyrazine-2-carboxamide (13 mg, 48% yield) as yellow solid. ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 10.10 (s, 1H), 8.85 (s, 2H), 8.30 (s, 1H), 8.02 (d, 2H), 7.90 (m, 2H), 7.40 (m, 1H), 7.02 (d, 2H), 3.80 (s, 3H), 2.82 (br s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_5\text{O}_3$: 378 (MH^+).

[0503] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[0504] 3-amino-N-methyl-6-{3-[(phenylcarbonyl)amino]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 10.60 (s, 1H), 8.72 (m, 2H), 8.30 (s, 1H), 8.05 (d, 2H), 7.92 (d, 1H), 7.82 (d, 1H), 7.60 (m, 3H), 7.45 (t, 1H), 2.86 (d, 3H); MS (EI) for $\text{C}_{19}\text{H}_{17}\text{N}_5\text{O}_2$: 348 (MH^+).

[0505] 3-amino-N-methyl-6-[3-({[3-(methyloxy)phenyl]carbonyl}amino)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 10.50 (s, 1H), 8.72 (br s, 2H), 8.30 (s, 1H), 7.92 (d, 1H), 7.85 (d, 1H), 7.50 (m, 6H), 7.20 (d, 1H), 3.90 (s, 3H), 2.90 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_5\text{O}_3$: 378 (MH^+).

[0506] 3-Amino-6-(3-({[4-chlorophenyl]carbonyl}amino)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 10.35 (s, 1H), 8.70 (br s, 2H), 8.25 (br s,

1H), 8.02 (br d, 2H), 7.92 (br d, 1H), 7.82 (br d, 1H), 7.65 (m, 2H), 7.40 (t, 1H), 2.92 (s, 3H); MS (EI) for C₁₉H₁₆N₅O₂Cl: 382 (MH⁺).

[0507] 3-Amino-N-methyl-6-(3-([(4-methylphenyl)carbonyl]amino)phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 10.20 (s, 1H), 8.70 (br s, 2H), 8.25 (s, 1H), 7.82 (m, 4H), 7.40 (t, 1H), 7.15 (d, 2H), 2.80 (d, 3H), 2.40 (s, 3H); MS (EI) for C₂₀H₁₉N₅O₂: 362 (MH⁺).

[0508] 3-Amino-6-{3-[(furan-2-ylcarbonyl)amino]phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 10.20 (s, 1H), 8.70 (br s, 2H), 8.24 (s, 1H), 7.95 (s, 1H), 7.92 (d, 1H), 7.80 (d, 1H), 7.40 (t, 1H), 7.26 (br s, 1H), 6.70 (br s, 1H), 2.80 (d, 3H); MS (EI) for C₁₇H₁₅N₅O₃: 338 (MH⁺).

[0509] 3-Amino-N-methyl-6-[3-([(3-(phenyloxy)phenyl)carbonyl]amino)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.75 (m, 2H), 8.28 (br s, 1H), 7.92 (d, 1H), 7.83 (br t, 2H), 7.64 (m, 1H), 7.58 (t, 1H), 7.45 (m, 4H), 7.26 (dd, 1H), 7.20 (br t, 1H), 7.12 (br d, 2H), 2.85 (d, 3H); MS (EI) for C₂₅H₂₁N₅O₃: 440 (MH⁺).

[0510] 3-Amino-6-{3-[(cyclopentylcarbonyl)amino]phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.92 (s, 1H), 8.70 (m, 2H), 8.10 (br s, 1H), 7.92 (d, 1H), 7.70 (d, 1H), 7.40 (t, 1H), 2.90 (m, 4H), 1.75 (br m, 8H); MS (EI) for C₁₈H₂₁N₅O₂: 340 (MH⁺).

[0511] 3-Amino-6-(3-([(4-cyanophenyl)carbonyl]amino)phenyl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 10.55 (br s, 1H), 8.70 (br s, 2H), 8.30 (br s, 1H), 8.15 (br d, 2H), 8.05 (br d, 2H), 7.92 (br d, 1H), 7.82 (br d, 1H), 7.40 (br t, 1H), 2.82 (br s, 3H); MS (EI) for C₂₀H₁₆N₆O₂: 373 (MH⁺).

Example 13

3-Amino-N-methyl-6-[3-([(phenylmethyl)amino]carbonyl)amino]phenyl]pyrazine-2-carboxamide

[0512] 3-amino-N-methyl-6-[3-([(phenylmethyl)amino]carbonyl)amino]phenyl]pyrazine-2-carboxamide: To a solution of 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide (0.25 g, 1.03 mmol) in dry tetrahydrofuran (2 mL) was added benzyl isocyanate (0.15 g, 1.12 mmol). The mixture was allowed to stir overnight. The solid product was filtered from the reaction mixture and triturated with hexane/ethyl

acetate (9:1). The solid was collected by filtration and dried to afford 3-amino-N-methyl-6-[3-({[(phenylmethyl)amino]carbonyl}amino)phenyl]pyrazine-2-carboxamide (0.26 g, 70% yield) as light yellow solid. ^1H NMR (400 MHz, d_6 -DMSO): 8.80 (m, 1H), 8.64 (s, 1H), 8.02 (s, 1H), 7.68 (d, 1H), 7.58 (d, 1H), 7.30 (m, 8H), 4.30 (d, 2H), 2.80 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{20}\text{N}_6\text{O}_2$: 377 (MH^+).

[0513] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[0514] 3-amino-6-[3-({[(4-fluorophenyl)amino]carbonyl}amino)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.80 (br s, 1H), 8.75 (br m, 1H), 8.70 (br s, 1H), 8.65 (br s, 1H), 7.92 (br s, 1H), 7.54 (br d, 1H), 7.45 (m, 2H), 7.39 (t, 1H), 7.12 (br t, 3H), 2.86 (d, 3H); MS (EI) for $\text{C}_{19}\text{H}_{17}\text{N}_6\text{O}_2\text{F}$: 381 (MH^+).

[0515] 3-amino-N-methyl-6-{3-[(4-(methyloxy)phenyl)amino]carbonyl}amino)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.78 (br m, 1H), 8.70 (br s, 1H), 8.59 (br d, 2H), 7.92 (br s, 1H), 7.72 (br d, 1H), 7.56 (br d, 1H), 7.34 (m, 3H), 6.82 (br d, 2H), 3.70 (s, 3H), 2.80 (br d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{20}\text{N}_6\text{O}_3$: 393 (MH^+).

[0516] According to the procedure described above, from 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide and the corresponding acids, the following compounds of the invention were prepared:

[0517] 3-amino-6-(3-[(4-chlorophenyl)acetyl]amino)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.30 (s, 1H), 8.70 (m, 2H), 8.10 (br s, 1H), 7.85 (d, 1H), 7.70 (d, 1H), 7.40 (m, 5H), 3.64 (s, 2H), 2.82 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{Cl}$: 396 (MH^+).

[0518] 3-amino-6-{3-[(diphenylacetyl)amino]phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.52 (s, 1H), 8.70 (m, 2H), 8.20 (br t, 1H), 7.85 (d, 1H), 7.72 (d, 1H), 7.32 (m, 11H), 5.20 (s, 1H), 2.82 (d, 3H); MS (EI) for $\text{C}_{26}\text{H}_{23}\text{N}_5\text{O}_2$: 438 (MH^+).

[0519] 3-amino-N-methyl-6-{3-[(phenylacetyl)amino]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.72 (m, 1H), 8.68 (s, 1H), 8.02 (br s, 1H), 7.82 (d, 1H), 7.72 (d, 1H), 7.34 (m, 6H), 7.28 (m, 1H), 3.70 (s, 2H), 2.80 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_5\text{O}_2$: 362 (MH^+).

- [0520] 3-amino-N-methyl-6-{3-[(3-phenylpropanoyl)amino]phenyl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.68 (m, 2H), 8.08 (br s, 1H), 7.84 (d, 1H), 7.68 (d, 1H), 7.40 (t, 1H), 7.28 (m, 5H), 7.20 (m, 1H), 2.92 (t, 2H), 2.82 (d, 3H), 2.62 (t, 2H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_2$: 376 (MH^+).
- [0521] 3-amino-6-(3-{[(2,4-dichlorophenyl)acetyl]amino}phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.30 (s, 1H), 8.62 (m, 2H), 8.10 (s, 1H), 7.82 (d, 1H), 7.62 (m, 3H), 7.40 (m, 4H), 3.92 (s, 2H), 2.82 (br s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{Cl}_2$: 430, 432 (MH^+).
- [0522] 3-amino-6-(3-{[(3,4-dichlorophenyl)acetyl]amino}phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.30 (s, 1H), 8.70 (m, 2H), 8.10 (s, 1H), 7.82 (d, 1H), 7.65 (m, 5H), 7.41 (m, 2H), 3.78 (s, 2H), 2.82 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{17}\text{N}_5\text{O}_2\text{Cl}_2$: 430, 432 (MH^+).
- [0523] 3-amino-N-methyl-6-[3-{[4-(trifluoromethyl)phenyl]acetyl}amino]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.40 (s, 1H), 8.80 (m, 2H), 8.10 (s, 1H), 7.90 (d, 1H), 7.70 (m, 4H), 7.60 (d, 2H), 7.40 (t, 1H), 3.80 (s, 2H), 2.84 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{18}\text{N}_5\text{O}_2\text{F}_3$: 430 (MH^+).
- [0524] 3-amino-6-{3-[(1,3-benzodioxol-5-ylacetyl)amino]phenyl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 10.20 (s, 1H), 8.70 (m, 2H), 8.10 (br s, 1H), 7.85 (d, 1H), 7.70 (d, 1H), 7.40 (t, 1H), 6.58 (m, 3H), 6.01 (s, 2H), 3.60 (s, 2H), 2.82 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_4$: 406 (MH^+).
- [0525] 9H-fluoren-9-ylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate: ^1H NMR (400 MHz, d_6 -DMSO): 9.73-9.67 (m, 1H), 8.68-8.61 (m, 2H), 8.00-7.20 (m, 2H), 8.00-7.96 (m, 1H), 7.90 (d, 2H), 7.86-7.26 (m, 9H), 4.49 (d, 2H), 4.32 (t, 1H), 2.84 (d, 3H); MS (EI) for $\text{C}_{27}\text{H}_{23}\text{N}_5\text{O}_3$: 466 (MH^+).
- [0526] phenylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate: ^1H NMR (400 MHz, d_6 -DMSO): 9.76 (s, 1H), 8.68-8.61 (m, 2H), 8.03-7.98 (m, 1H), 8.00-7.00 (m, 2H), 7.82-7.76 (m, 1H), 7.54-7.47 (m, 1H), 7.46-7.30 (m, 4H), 5.18-5.14 (m, 2H), 4.50 (d, 2H), 2.87-2.82 (m, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_5\text{O}_3$: 378 (MH^+).
- [0527] (1R,2S,5R)-5-methyl-2-(1-methylethyl)cyclohexyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate: ^1H NMR (400 MHz, d_6 -DMSO): 9.56 (s, 1H), 8.67-8.61 (m, 2H), 8.01-7.99 (m, 1H), 7.90-7.20 (m, 2H), 7.77 (d,

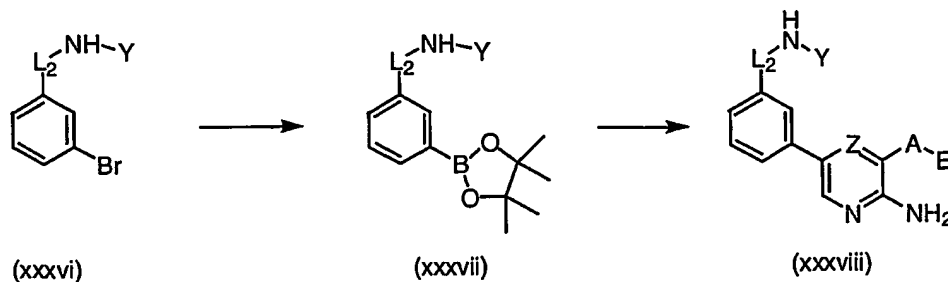
1H), 7.54-7.49 (m, 1H), 7.37-7.32 (m, 1H), 4.56 (td, 1H), 2.85 (d, 3H), 1.72-0.68 (m, 18H); MS (EI) for $C_{23}H_{31}N_5O_3$: 426 (MH^+).

[0528] 1,1-dimethylethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate: 1H NMR (400 MHz, d_6 -DMSO): 9.29 (s, 1H), 8.68-8.61 (m, 2H), 7.99-7.96 (m, 1H), 8.00-7.20 (br, 2H), 7.77-7.72 (m, 1H), 7.52-7.47 (m, 1H), 7.35-7.30 (m, 1H), 2.83 (d, 3H), 1.47 (s, 9H); MS (EI) for $C_{17}H_{21}N_5O_3$: 344 (MH^+).

Example 14

[0529] Scheme 8 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene or pyridylene, X is $-SO_2NH-$ or $-C(=O)NH-$ wherein the phenylene or pyridylene is attached to the sulfur or the carbon atom of X, and Y is attached to the nitrogen of X. As described above, W, X, and Y all are optionally substituted, but for simplicity this is not shown here. As depicted, intermediate (xxxvi) is incorporated into a compound of the invention, (xxxviii), where Z is N or C, and L_2 is a substructure of X. In Scheme 8, L_2 represents either $-SO_2-$ or $-C(=O)-$. The sequence of transformations regarding such intermediates may vary depending on the nature of each of W, X, and Y. For example, when L_2 is $-SO_2-$, the sulfonamide is typically generated prior to the conversion of the bromo group of (xxxvi) to the boronic ester group in intermediate (xxxvii). The same is true when L_2 is $-C(=O)-$ and W is pyridylene. In cases where W is a substituted benzene, L_2 is $-C(=O)-$, and Y is an alkyl group, the amide bond is formed after the formation of (xxxviii) using synthetic steps known in the art. Conversion of (xxxvi) to (xxxvii) is carried out using bis(pinacolato)diboron based on procedures known to those skilled in the art. The transformation of (xxxvii) to (xxxviii) is accomplished using Suzuki coupling conditions, for example.

Scheme 8



3-Amino-6-(4-chloro-3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

- [0530] Methyl 2-chloro-5-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)benzoate: To a solution of methyl 5-bromo-2-chlorobenzoate (0.25 g, 1.01 mmol) in anhydrous dimethylsulfoxide (6 mL) were added bis(pinacolato)diboron (0.28 g, 1.10 mmol), potassium acetate (0.29 g, 3.01 mmol), and [1,1'-bis (diphenylphosphino)ferrocene] dichloropalladium(II) complex with dichloromethane (1:1, 25 mg, 0.03 mmol). The solution was degassed by passing nitrogen gas for 3-5 min and heated at 80-85° C for 3 h. Reaction cooled to room temperature and diluted with ethyl acetate (250 mL). The organic layer was washed with saturated aqueous sodium bicarbonate (2 X 40 mL) and saturated aqueous sodium chloride (50 mL), dried over magnesium sulfate, filtered, and concentrated to afford methyl 2-chloro-5-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)benzoate (0.12 g, 42% yield) as semi solid. ¹H NMR (400 MHz, CDCl₃): 8.20 (br d, 1H), 7.80 (dd, 1H), 7.43 (d, 1H), 3.92 (s, 3H), 1.25 (s, 12H); MS (EI) for C₁₄H₁₈N₄O₂BCl: 297 (MH⁺).
- [0531] Methyl 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-2-chlorobenzoate: To a solution of methyl 2-chloro-5-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)benzoate (0.16 g, 0.55 mmol) in toluene (2 mL)) were added 3-amino-6-bromo-N-methyl-2-pyrazincarboxamide (0.13 g, 0.58 mmol), tetrakis(triphenylphosphin)palladium(0) (64 mg, 0.06 mmol), ethanol (2 mL), and 2.0 N sodium carbonate (0.42 g, 4.00 mmol). The solution was degassed with nitrogen gas for 3-5 min and heated at 90-100° C for 2 h. Reaction cooled to room temperature and diluted with ethyl acetate (200 mL). The organic layer was washed water (50 mL) and saturated aqueous sodium chloride (40 mL), dried over magnesium sulfate, filtered, and concentrated to afford crude product. Trituration with methanol (1 mL), filtration and drying gave methyl 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-2-chlorobenzoate (96 mg, 56% yield) as yellow solid. ¹H NMR (400 MHz, CDCl₃): 8.62 (br s, 1H), 8.28 (d, 1H), 7.92 (dd, 2H), 7.52 (d, 1H), 4.02 (s, 3H), 3.02 (s, 3H); MS (EI) for C₁₄H₁₃N₄O₂Cl: 321 (MH⁺).
- [0532] 5-{5-Amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-2-chlorobenzoic acid: To a solution of methyl 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-2-chlorobenzoate (96 mg, 31 mmol) in methanol/tetrahydrofuran (1:1, 5.0 mL) was added 2.0 N aqueous sodium hydroxide (0.16 g, 4.00 mmol). The reaction was stirred overnight

at room temperature. Reaction was neutralized with 2.0 N aqueous hydrochloric acid (4 mL) and diluted with ethyl acetate (100 mL). The organic layer was washed with water (20 mL), saturated aqueous sodium chloride (30 mL), dried over magnesium sulfate, filtered, and concentrated to afford 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-2-chlorobenzoic acid (45 mg, 48% yield). ^1H NMR (400 MHz, d_6 -DMSO): 13.42 (br s, 1H), 8.90 (br s, 1H), 8.45 (d, 1H), 8.32 (dd, 1H), 7.85 (br s, 2H), 7.60 (d, 2H), 2.82 (s, 3H); MS (EI) for $\text{C}_{13}\text{H}_{11}\text{N}_4\text{O}_3\text{Cl}$: 307 (MH^+).

[0533] (d)-3-Amino-6-(4-chloro-3-[[[(phenylmethyl)amino]carbonyl]phenyl])-N-methylpyrazine-2-carboxamide: The title compound was synthesized according to the procedure described above from 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-2-chlorobenzoic acid and benzylamine. Yield 99%. ^1H NMR (400 MHz, d_6 -DMSO): 9.10 (t, 1H), 8.90 (s, 2H), 8.26 (m, 2H), 7.60 (d, 1H), 7.40 (m, 4H), 7.14 (t, 1H), 4.50 (d, 2H), 2.82 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{Cl}$: 396, 398 (MH^+).

[0534] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared.

[0535] 3-amino-6-(4-fluoro-3-[[[(phenylmethyl)amino]carbonyl]phenyl])-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.64 (m, 2H), 7.98 (m, 2H), 7.40 (m, 3H), 7.32 (m, 1H), 7.20 (m, 2H), 7.15 (m, 1H), 4.70 (d, 2H), 3.02 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{F}$: 380 (MH^+).

[0536] 3-amino-N-methyl-6-(4-(methyloxy)-3-[[[(phenylmethyl)amino]carbonyl]phenyl])pyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.80 (s, 1H), 8.60 (s, 1H), 8.20 (br s, 1H), 7.90 (m, 2H), 7.30 (m, 5H), 7.20 (d, 1H), 4.70 (d, 2H), 3.92 (s, 3H), 3.02 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}_3$: 392 (MH^+).

[0537] 3-amino-6-(2-fluoro-5-[[[(phenylmethyl)amino]carbonyl]phenyl])-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): 8.62 (d, 1H), 8.30 (dd, 1H), 7.90 (br s, 1H), 7.70 (m, 1H), 7.30 (m, 4H), 7.26 (s, 1H), 7.20 (t, 1H), 6.50 (br s, 1H), 4.65 (d, 2H), 3.02 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{F}$: 380 (MH^+).

Example 15

3-Amino-N-methyl-6-(3-[[[(phenylmethyl)amino]sulfonyl]phenyl])pyrazine-2-carboxamide

[0538] 3-Bromo-N-(phenylmethyl)benzenesulfonamide: To a solution of 3-bromosulfonyl chloride (1.31 g, 5.14 mmol) in dry tetrahydrofuran (10 mL) at 0° C were added triethylamine (0.62 g, 6.17 mmol) and benzylamine (0.50 g, 4.57 mmol). The reaction was slowly warmed to room temperature and stirred overnight. Reaction was treated with 2.0 N aqueous hydrochloric acid (10 mL) and diluted with ethyl acetate (300 mL). The organic layer was washed with water (30 mL), saturated aqueous sodium bicarbonate (40 mL), saturated aqueous sodium chloride (40 mL), dried over magnesium sulfate, filtered, and concentrated to afford 3-Bromo-N-(phenylmethyl)benzenesulfonamide (1.32 g, 78% yield). ¹H NMR (400 MHz, CDCl₃): 7.34 (t, 1H), 7.85 (s, 1H), 7.80 (m, 2H), 7.50 (t, 1H), 7.22 (m, 5H), 4.02 (d, 2H); MS (EI) for C₁₃H₁₂NO₂SBr: 326 (MH⁺).

[0539] N-(phenylmethyl)-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)benzenesulfonamide: The title compounds was synthesized according to the procedure described above. Yield 69 %. ¹H NMR (400 MHz, CDCl₃): 8.25 (s, 1H), 7.92 (m, 2H), 7.43 (t, 1H), 7.23 (m, 6H), 4.12 (d, 2H), 1.23 (s, 12H); MS (EI) for C₁₉H₂₄NO₄SB: 374 (MH⁺).

[0540] 3-Amino-N-methyl-6-(3-([(phenylmethyl)amino]sulfonyl)phenyl)pyrazine-2-carboxamide was synthesized from N-(phenylmethyl)-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)benzenesulfonamide and 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide as yellow solid. Yield 20%. ¹H NMR (400 MHz, d₆-DMSO): 8.84 (s, 1H), 8.80 (m, 1H), 8.46 (d, 1H), 8.38 (br s, 1H), 8.20 (t, 1H), 7.78 (d, 1H), 7.66 (t, 1H), 7.26 (m, 4H), 7.18 (m, 1H), 4.02 (d, 2H), 2.82 (d, 3H); MS (EI) for C₁₉H₁₉N₅O₃S: 398 (MH⁺).

Example 16

3-Amino-N-methyl-6-(5-([(phenylmethyl)amino]carbonyl)pyridin-3-yl)pyrazine-2-carboxamide

[0541] 5-bromo-N-(phenylmethyl)pyridine-3-carboxamide was synthesized according to the procedure described above from 5-bromonicotinic acid and benzylamine. Yield 90%. ¹H NMR (400 MHz, CDCl₃): 8.83 (d, 1H), 8.72 (d, 1H), 8.25 (t, 1H), 7.30 (m, 5H), 6.58 (br s, 1H), 4.62 (d, 2H); MS (EI) for C₁₃H₁₁N₂OBr: 291 (MH⁺).

- [0542] N-(phenylmethyl)-5-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)pyridine-3-carboxamide was synthesized according to the procedure above from 5-bromo-N-(phenylmethyl)pyridine-3-carboxamide and bis(pinacolato)diboron. Yield 49%. ¹H NMR (400 MHz, CDCl₃): 9.15 (br s, 1H), 9.02 (br s, 1H), 8.35 (br s, 1H), 7.32 (m, 5H), 6.42 (br t, 1H), 4.62 (d, 2H), 1.29 (s, 12H); MS (EI) for C₁₉H₂₃N₂O₃B: 339 (MH⁺).
- [0543] 3-amino-N-methyl-6-(5-[(phenylmethyl)amino]carbonyl)pyridin-3-yl)pyrazine-2-carboxamide was synthesized according to the procedure described above from N-(phenylmethyl)-5-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)pyridine-3-carboxamide and 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide. Yield 38%. ¹H NMR (400 MHz, d₆-DMSO): 9.58 (d, 1H), 9.36 (t, 1H), 9.02 (m, 3H), 8.86 (t, 1H), 7.34 (m, 4H), 7.24 (m, 1H), 4.50 (d, 2H), 2.83 (d, 3H); MS (EI) for C₁₉H₁₈N₆O₂: 363 (MH⁺).
- [0544] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:
- [0545] 3-amino-6-[5-([(2,6-difluorophenyl)methyl]amino)carbonyl]pyridin-3-yl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.54(d, 1H), 9.12 (t, 1H), 8.92 (m, 3H), 8.79 (t, 1H), 7.42 (m, 1H), 7.12 (t, 2H), 4.60 (d, 2H), 2.82 (d, 3H); MS (EI) for C₁₉H₁₆N₆O₂F₂: 399 (MH⁺).
- [0546] 3-amino-6-(5-[(biphenyl-2-ylmethyl)amino]carbonyl)pyridin-3-yl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.52 (d, 2H), 9.15 (t, 1H), 8.92 (m, 3H), 8.80 (t, 1H), 7.43 (m, 8H), 7.24 (br d, 1H), 4.50 (d, 2H), 2.82 (d, 3H); MS (EI) for C₂₅H₂₂N₆O₂: 439 (MH⁺).
- [0547] 3-amino-6-(5-[(2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino]carbonyl)pyridin-3-yl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.52 (br t, 2H), 9.25 (t, 1H), 6.95 (t, 1H), 8.92 (m, 1H), 8.82 (m, 2H), 8.80 (m, 1H), 6.82 (br s, 2H), 4.40 (br s, 2H), 4.20 (s, 4H), 2.83 (d, 3H); MS (EI) for C₂₁H₂₀N₆O₄: 421 (MH⁺).

Example 17

3-Amino-N-methyl-6-[3-[(2-phenylethyl)amino]phenyl]pyrazine-2-carboxamide

- [0548] 3-bromo-N-(2-phenylethyl)aniline: To a solution of 3-bromoaniline (3.21 g, 18.60 mmol) in dry tetrahydrofuran (45 mL) were added triethylamine (2.82 g, 27.91 mmol) and phenylacetyl chloride (93.45 g, 22.30 mmol) at 0° C. The reaction warmed to room temperature and stirred for 4 h. The reaction mixture was treated with 2.0 N aqueous hydrochloric acid (20 mL) at 0° C and diluted with ethyl acetate (300 mL). The organic layer was washed with water (20 mL), saturated aqueous sodium bicarbonate (50 mL) and saturated aqueous sodium chloride (50 mL), dried over magnesium sulfate, filtered, and concentrated to afford crude N-(3-bromophenyl)-2-phenylacetamide (4.84 g, 86% yield) which was used in the next step without any further purification.
- [0549] To a solution of crude N-(3-bromophenyl)-2-phenylacetamide (3.02 g, 10.4 mmol) in dry tetrahydrofuran (25 mL) was added 1.0 N borane-tetrahydrofuran complex (1:1, 26 mL, 26.12 mmol) from an addition funnel at 0° C. The reaction was refluxed under nitrogen overnight. Reaction mixture was cooled to room temperature, treated with 2.0 N aqueous sodium hydroxide (30 mL) and then stirred for 20 min. The reaction was then cooled again at 0° C and neutralized with 2.0 N aqueous hydrochloric acid (60 mL) and further stirred for 30 min. The aqueous layer was extracted with ethyl acetate (3X100 mL). The organic layer was washed with water (20 mL), saturated aqueous sodium chloride (50 mL), dried over magnesium sulfate, filtered, and concentrated to afford 3-Bromo-N-(2-phenylethyl)aniline (2.55 g, 85% yield) as brown oil with reasonable purity: ¹H NMR (400 MHz, CDCl₃): 7.30 (t, 2H), 7.20 (m, 3H), 6.95 (t, 1H), 6.78 (br d, 1H), 6.70 (t, 1H), 6.45 (dd, 1H), 3.62 (br s, 1H), 3.40 (m, 2H), 2.92 (t, 2H); MS (EI) for C₁₄H₁₂NOBr: 290 (MH⁺).
- [0550] N-(2-phenylethyl)-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)aniline was synthesized according to the procedure described above from 3-bromo-N-(2-phenylethyl)aniline and bis(pinacolato)diboron. Yield 72%. ¹H NMR (400 MHz, CDCl₃): 7.80 (m, 2H), 7.18 (m, 5H), 7.20 (br d, 1H), 6.70 (br t, 1H), 3.40 (t, 2H), 2.85 (t, 2H), 1.23 (s, 12H); MS (EI) for C₂₀H₂₆NO₂B: 324 (MH⁺).
- [0551] 3-Amino-N-methyl-6-{3-[(2-phenylethyl)amino]phenyl}pyrazine-2-carboxamide was synthesized according to the procedure described above from N-(2-phenylethyl)-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)aniline and 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide. Yield 39%. ¹H NMR (400 MHz, CDCl₃): 8.58 (s, 1H), 8.02 (br s, 1H), 7.32 (m, 3H), 7.26 (m, 3H), 7.18 (br d, 1H), 7.02 (br t, 1H), 6.64 (dd, 1H), 3.82 (br s, 1H), 3.51 (t, 2H), 3.02 (d, 3H), 2.94 (t, 2H); MS (EI) for C₂₀H₂₁N₅O: 348 (MH⁺).

[0552] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[0553] 3-amino-N-methyl-6-{1-[(phenylmethyl)amino]isoquinolin-7-yl}pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 11.15 (m, 1H), 9.64 (s, 1H), 9.30 (m, 1H), 9.15 (s, 1H), 8.84 (m, 1H), 8.03 (m, 1H), 7.58 (m, 1H), 7.50 (m, 2H), 7.37 (m, 2H), 7.28 (m, 2H), 5.08 (br s, 2H), 4.98 (d, 2H), 2.87 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{20}\text{N}_6\text{O}$: 385.2 (MH^+).

[0554] 3-amino-6-{1-[(1S)-2,3-dihydro-1H-inden-1-ylamino]isoquinolin-7-yl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.07 (s, 1H), 8.89 (m, 2H), 8.59 (d, 1H), 7.90 (d, 1H), 7.79 (d, 2H), 7.66 (br s, 2H), 7.28 (m, 1H), 7.25 (m, 1H), 7.21 (t, 1H), 7.14 (t, 1H), 6.97 (d, 1H), 6.02 (m, 1H), 3.04 (m, 1H), 2.91 (m, 1H), 2.85 (d, 3H), 2.61 (m, 1H), 2.07 (m, 1H); MS (EI) for $\text{C}_{24}\text{H}_{22}\text{N}_6\text{O}$: 411.2 (MH^+).

[0555] 3-amino-6-{3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]-2,3-dihydro-1H-inden-5-yl}-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.77 (s, 1H), 8.73 (m, 1H), 8.01 (m, 2H), 7.57 (br s, 2H), 7.48 (m, 1H), 7.37 (m, 1H), 7.29 (d, 1H), 7.21 (m, 1H), 7.15 (m, 2H), 4.34 (m, 2H), 2.97 (m, 2H), 2.84 (d, 3H), 2.77 (m, 2H), 2.44 (m, 1H), 2.34 (m, 1H), 1.83 (m, 2H); MS (EI) for $\text{C}_{24}\text{H}_{25}\text{N}_5\text{O}$: 400.2 (MH^+).

[0556] 3-amino-N-methyl-6-[3-([2-(phenylamino)ethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide was synthesized according to the procedure described in above from N-phenylethane-1,2-diamine and 3-{5-amino-6-[(methylamino)carbonyl]pyrazine-2-yl}benzoic acid. Yield 45%: ^1H NMR (400 MHz, CDCl_3): 8.90 (s, 1H), 8.82 (br d, 1H), 8.70 (br t, 1H), 8.45 (br s, 1H), 8.32 (br d, 1H), 7.82 (br d, 1H), 7.52 (t, 1H), 7.02 (t, 2H), 6.62 (d, 2H), 6.50 (t, 1H), 5.72 (br t, 1H), 3.52 (d, 2H), 3.20 (d, 2H), 2.92 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{22}\text{N}_6\text{O}_2$: 391 (MH^+).

[0557] 6,6'-[(Oxomethanediyl)bis(iminobenzene-3,1-diyl)]bis(3-amino-N-methylpyrazine-2-carboxamide): To a solution of 3-Amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide (0.10 g, 0.41 mmol) in dry tetrahydrofuran were added triethylamine (46 mg, 0.45 mmol) and 4-nitrophenyl chloroformate (91 mg, 0.45 mmol). The reaction of was allowed to stir overnight. A yellow solid material from the crude reaction was filtered and washed with diethyl ether (50 mL) and hexanes (50 mL) to yield 6,6'-[(oxomethanediyl)bis(iminobenzene-3,1-diyl)]bis(3-amino-N-methylpyrazine-2-

carboxamide (0.11 g, 54% yield): ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 9.25 (br s, 2H), 8.75 (br d, 2H), 8.70 (s, 2H), 8.01 (s, 2H), 7.74 (br d, 2H), 7.60 (br d, 2H), 7.32 (t, 2H), 6.82 (br d, 2H), 2.80 (br s, 6H); MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_{10}\text{O}_3$: 513 (MH^+).

[0558] 3-Amino-6-biphenyl-3-yl-N-methylpyrazine-2-carboxamide was synthesized according to the procedure described in above. Yield 30%. ^1H NMR (400 MHz, CDCl_3): 8.65 (br s, 1H), 8.03 (m, 1H), 8.01 (br s, 1H), 7.82 (dd, 1H), 7.65 (m, 3H), 7.50 (m, 3H), 7.40 (t, 1H), 3.02 (br s, 3H); MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_4\text{O}$: 305 (MH^+)

[0559] Ethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate was synthesized according to the procedure described in above from 3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide and ethyl chloroformate. Yield 55%: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 9.60 (br s, 1H), 8.68 (br s, 2H), 8.12 (br s, 1H), 7.80 (d, 1H), 7.52 (d, 1H), 7.39 (t, 1H), 4.06 (q, 2H), 2.85 (br s, 3H), 1.24 (t, 3H); MS (EI) for $\text{C}_{15}\text{H}_{17}\text{N}_5\text{O}_3$: 316 (MH^+)

[0560] 3-Amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide. To a solution of 3-amino-6-bromo-N-methyl-2-pyrazinecarboxamide (1.23 g, 5.30 mmol) in N,N-dimethylformamide (30 mL) were added [1,1'-bis(diphenylphosphino)ferrocene]dichloropalladium(II) complex with dichloromethane (1:1, 433 mg, 0.53mmol), 3-aminophenylboronic acid (1.15 g, 7.41 mmol), and triethylamine (1.07 g, 10.6mmol). The solution was degassed with nitrogen 3-5 min. The reaction was heated at 85-90° C for 15 h with stirring. The reaction was allowed to cool to room temperature and diluted with ethyl acetate (400 mL). The organic layer was washed with saturated aqueous sodium bicarbonate (50 mL), saturated aqueous sodium chloride (50 mL), dried over magnesium sulfate, filtered, and concentrated to give crude product. To the crude product was added 4.0 N hydrochloric acid (1,4-dioxane solution, 30 mL). The resulting solid was filtered and washed with diethyl ether/methanol (4:1). The solid was dissolved in methanol and neutralized with sufficient AG 1-X8 Resin (hydroxide form) that the pH of the solution became basic. The mixture was filtered, and the filtrate was concentrated on a rotary evaporator at reduced pressure to afford 3-Amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide (0.70 g, 54% yield) as dark green solid. ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 8.70(br s, 1H), 8.62 (s, 1H), 7.60 (br s, 2H), 7.25 (m, 2H), 7.10 (t, 1H), 6.15 (d, 1H), 5.06 (s, 2H), 2.8 (br s, 3H); MS (EI) for $\text{C}_{12}\text{H}_{13}\text{N}_5\text{O}$: 244 (MH^+).

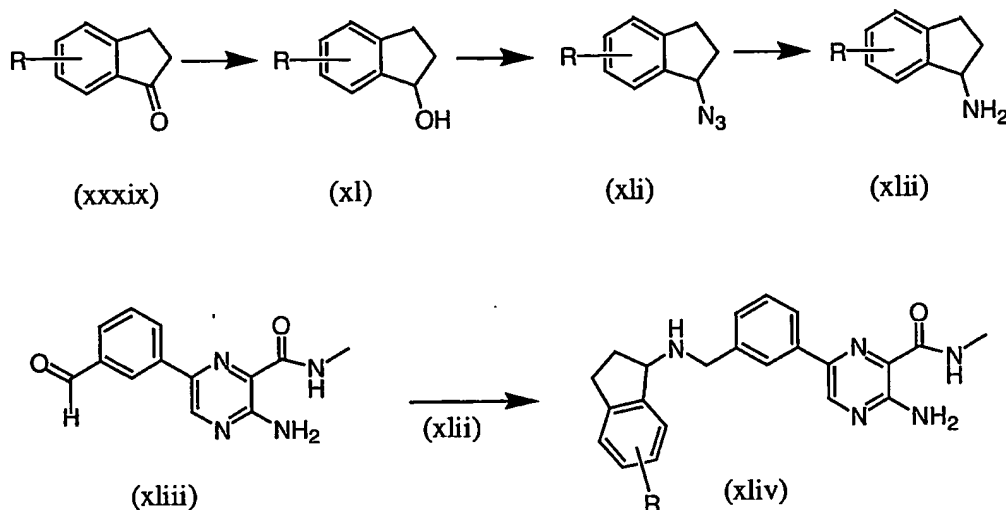
Example 18

[0561] Scheme 9 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene, X is -CH₂NH-, and Y is an indanyl group. Note, as defined above any of W, X, or Y are optionally substituted; this is only a set of examples. As depicted, intermediate (xxxix) is incorporated into a compound of the invention, (xliv). For example, optionally substituted indanone (xxxix) was to the corresponding alcohol (xl) using sodium borohydride or other appropriate reducing agent, depending upon, among other things, the substituent, R. The alcohol group in (xl) is transformed into an azide group in (xli) using diphenylphosphoryl azide and 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU), for example. Reduction of azide (xli) is carried out typically, but not necessarily, using tin chloride give amine (xl ii). Amine (xl ii) is coupled to aldehyde (xl iii), via reductive amination to afford (xl iv). Amine (xl ii) could also be coupled to a benzyl bromide, analogous to aldehyde (xl iii) to form (xl iv), for example.

3-amino-N-methyl-6-[3-([5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]pyrazine-2-carboxamide

[0562] 5-(2-thienyl)-2,3-dihydro-1H-inden-1-one. To a solution of 5-bromo-1-indanone (0.52g, 2.5 mmol) in toluene (5 mL) were added tetrakis(triphenylphosphine)-palladium(0) (29 mg, 0.03 mmol), thiophene-2-boronic acid (0.48 g, 3.7 mmol), ethanol (5 mL), and 2.0 N sodium carbonate (3 mL). The solution was flushed with nitrogen for 3 minutes. The reaction was heated at 110-120° C for 4h. Reaction cooled to room temperature and diluted with ethyl acetate (200 mL). The organic layer was washed with saturated aqueous sodium bicarbonate (2X50 mL), water (40 mL), saturated aqueous sodium chloride (50 mL), dried over magnesium sulfate, filtered and concentrated to afford 5-(2-thienyl)-2,3-dihydro-1H-inden-1-one (0.50g, 94% yield) as pure solid product. ¹H NMR (400 MHz, CDCl₃) 7.80 (d, 1H), 7.70 (s, 1H), 7.62 (d, 1H), 7.43 (dd, 1H), 7.38 (dd, 1H), 7.12 (m, 1H), 3.20 (m, 2H), 2.66 (m, 2H).

Scheme 9



[0563] 5-(2-thienyl)-2,3-dihydro-1H-inden-1-ol: To a slurry of sodium borohydride (0.22 g, 5.86 mmol) in methanol (6mL) at 0° C was added a solution of 5-(2-thienyl)-2,3-dihydro-1H-inden-1-one (0.50 g, 2.34 mmol) in methanol (2 mL). The reaction was slowly warmed to room temperature and stirred until the reaction was completed as monitored by thin layer chromatography (4 h). The reaction mixture was carefully quenched with 1.0 N aqueous hydrochloric acid at 0° C until a clear solution was obtained. Diluted with ethyl acetate (200 mL), washed with saturated aqueous sodium bicarbonate, saturated aqueous sodium chloride (brine), and dried over anhydrous magnesium sulfate. Filtration, concentration and column chromatography on silica (7:3 hexanes/ethyl acetate) gave 5-(2-thienyl)-2,3-dihydro-1H-inden-1-ol (0.30 g, 59% yield) as solid. ¹H NMR (400 MHz, CDCl₃): 7.48 (m, 2H), 7.38 (br d, 1H), 7.26 (m, 2H), 7.08 (dd, 1H), 5.21 (t, 1H), 3.04 (m, 1H), 2.80 (m, 1H), 2.50 (m, 1H), 2.10 (br s, 1H), 1.93 (m, 1H).

[0564] 2-(1-azido-2,3-dihydro-1H-indene-5-yl)thiophene: To a solution of 5-(2-thienyl)-2,3-dihydro-1H-inden-1-ol (0.30 g, 1.39 mmol) in tetrahydrofuran (6 mL) was added diphenylphosphoryl azide (0.58 g, 2.10 mmol) at 0° C. The reaction was stirred for 5 min prior to the addition of 1,8-diazabicyclo[5.4.0]undec-7-ene (DBU, 0.32 g, 2.10 mmol). The reaction was further stirred for 15 min at 0° C before the reaction was allowed to warm to room temperature. The reaction was stopped after 1h when the reaction was completed as monitored by thin layer chromatography. Diluted with ethyl acetate (200 mL), washed with water, saturated aqueous sodium chloride (brine), and dried over

anhydrous magnesium sulfate. Filtration, concentration and column chromatography on silica (7:3 hexanes/ethyl acetate) gave 2-(1-azido-2,3-dihydro-1H-indene-5-yl)thiophene (0.24g, 72% yield) as oil. ¹H NMR (400 MHz, CDCl₃) 7.54 (m, 2H), 7.40 (m, 2H), 7.24 (m, 1H), 7.08 (m, 1H), 5.85 (t, 1H), 3.12 (m, 1H), 2.90 (m, 1H), 2.45 (m, 1H), 2.19 (m, 1H).

[0565] [5-(2-thienyl)-2,3-dihydro-1H-indene-1-yl]amine: To a solution of 2-(1-azido-2,3-dihydro-1H-indene-5-yl)thiophene (0.24 g, 0.99 mmol) in methanol (2mL) was added solid tin chloride dihydrate (0.45 g, 1.99 mmol) at room temperature under nitrogen. The reaction was stirred at room temperature overnight. The crude reaction mixture was poured into a separatory funnel and diluted with ethyl acetate (250mL). The organic layer was washed with 2.0 N aqueous sodium hydroxide (50mL), saturated aqueous sodium chloride (brine), and dried over anhydrous magnesium sulfate. Filtration, concentration and column chromatography on silica (9:1 dichloromethane/methanol) gave [5-(2-thienyl)-2,3-dihydro-1H-indene-1-yl]amine (0.15 g, 72% yield) as semi-solid: ¹H NMR (400 MHz, CDCl₃) 7.54 (m, 1H), 7.40 (d, 2H), 7.29 (m, 2H), 7.08 (m, 1H), 4.40 (t, 1H), 3.05 (m, 1H), 2.90 (m, 1H), 2.56 (m, 1H), 1.80 (m, 1H).

[0566] 3-amino-N-methyl-6-[3-([5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]pyrazine-2-carboxamide: To a solution of 5-(2-thienyl)-2,3-dihydro-1H-indene-1-yl]amine (73 mg, 0.34 mmol) in acetonitril (3 mL) were added 3-amino-6-[3-(bromomethyl)phenyl]-N-methylpyrazine-2-carboxamide (41 mg, 0.13 mmol) and N,N-diisopropylethylamine (22 mg, 0.17 mmol) at room temperature. The reaction was stirred overnight. Diluted with ethyl acetate (100 mL) and washed with water, saturated aqueous sodium chloride (brine), and dried over anhydrous magnesium sulfate. Filtration and concentration and column purification on silica (9:1 ethyl acetate/methanol) gave 3-amino-N-methyl-6-[3-([5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]pyrazine-2-carboxamide (30 mg, 51% yield) as yellow solid. ¹NMR (400 MHz, CD₃OD): 8.70 (s, 1H), 8.15 (br s, 1H), 7.86 (d, 1H), 7.49 (m, 6H), 7.32 (m, 2H), 7.05 (m, 1H), 4.35 (m, 1H), 3.92 (d, 2H), 3.10 (m, 4H), 2.95 (s, 3H), 2.85 (m, 1H), 2.45 (m, 1H), 2.10 (m, 1H); MS (EI) for C₂₆H₂₅N₅OS: 456 (MH⁺).

[0567] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

- [0568] 3-amino-6-(3-{[(5-bromo-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, d₆-DMSO): 9.80 (br s, 2H), 9.40 (br s, 1H), 8.84 (s, 1H), 8.80 (s, 1H), 8.25 (d, 1H), 7.88 (br s, 1H), 7.80 (d, 1H), 7.48 (m, 4H), 4.82 (s, 1H), 4.24 (s, 2H), 3.25 (s, 3H), 3.20 (m, 1H), 2.80 (m, 1H), 2.50 (m, 1H), 2.40 (m, 1H); MS (EI) for C₂₂H₂₂N₅OBr: 453, 455 (MH⁺).
- [0569] 3-amino-6-(3-{[(5-furan-2-yl-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, CD₃OD): 8.65 (s, 1H), 8.15 (s, 1H), 7.88 (d, 1H), 7.50 (m, 6H), 6.70 (d, 1H), 6.45 (m, 1H), 4.40 (m, 1H), 4.01 (d, 2H), 3.15 (m, 3H), 2.96 (s, 3H), 2.85 (m, 1H), 2.50 (m, 1H), 2.45 (m, 1H), 2.10 (m, 1H); MS (EI) for C₂₆H₂₅N₅O₂: 440 (MH⁺).
- [0570] 3-amino-6-(3-{[(5-chloro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.58 (s, 1H), 8.20 (br s, 1H), 8.08 (br s, 1H), 7.70 (d, 1H), 7.40 (m, 3H), 7.20 (s, 1H), 7.15 (d, 1H), 4.40 (m, 1H), 3.80 (s, 2H), 3.10 (m, 4H), 2.80 (s, 1H), 2.45 (m, 1H), 2.20 (m, 1H); MS (EI) for C₂₂H₂₂N₅OCl: 408 (MH⁺).
- [0571] 3-amino-6-(3-{[(5-furan-3-yl-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.62 (br s, 1H), 8.05 (br s, 1H), 7.89 (br s, 1H), 7.78 (m, 2H), 7.40 (m, 6H), 6.70 (br s, 1H), 4.35 (m, 1H), 4.01 (d, 1H), 3.10 (m, 4H), 2.95 (s, 3H), 2.85 (m, 1H), 2.50 (m, 1H), 1.85 (m, 1H); MS (EI) for C₂₆H₂₅N₅O₂: 440 (MH⁺).
- [0572] 3-amino-N-methyl-6-[3-({[5-(3-thienyl)-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]pyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.64 (s, 1H), 8.02 (br s, 1H), 7.90 (br s, 1H), 7.78 (m, 1H), 7.44 (m, 6H), 7.38 (m, 2H), 4.40 (m, 1H), 4.02 (d, 2H), 3.15 (m, 4H), 2.90 (m, 1H), 2.50 (m, 1H), 1.90 (m, 1H); MS (EI) for C₂₆H₂₅N₅OS: 456 (MH⁺).
- [0573] 3-amino-N-methyl-6-(3-{[(5-phenyl-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)pyrazine-2-carboxamide: ¹NMR (400 MHz, d₆-DMSO): 8.82 (m, 2H), 8.16 (br s, 1H), 8.05 (d, 1H), 7.62 (d, 2H), 7.45 (m, 7H), 7.34 (m, 1H), 4.24 (m, 1H), 3.90 (m, 2H), 3.01 (m, 1H), 2.80 (m, 4H), 2.38 (m, 1H), 1.90 (m, 1H); MS (EI) for C₂₈H₂₇N₅O: 450 (MH⁺).
- [0574] 3-amino-N-methyl-6-[3-({[6-(2-thienyl)-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]pyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.60 (s,

1H), 7.99 (br s, 1H), 7.88 (br s, 1H), 7.75 (m, 1H), 7.60 (s, 1H), 7.45 (m, 3H), 7.24 (m, 3H), 7.04 (m, 1H), 4.40 (m, 1H), 4.02 (d, 2H), 3.08 (m, 4H), 2.83 (m, 1H), 2.50 (m, 1H), 1.86 (m, 1H); MS (EI) for C₂₆H₂₅N₅OS: 456 (MH⁺).

[0575] 3-amino-N-methyl-6-[3-([5-(4-methyl-2-thienyl)-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]pyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.60 (s, 1H), 8.01 (br s, 1H), 7.88 (br s, 1H), 7.74 (m, 1H), 7.60 (s, 1H), 7.40 (m, 5H), 7.08 (br s, 1H), 6.80 (br s, 1H), 4.70 (m, 1H), 4.02 (d, 2H), 3.06 (m, 4H), 2.85 (m, 1H), 2.50 (s, 3H), 1.90 (m, 1H); MS (EI) for C₂₇H₂₇N₅OS: 470 (MH⁺).

[0576] 3-amino-6-{3-[(2,3-dihydro-1-benzofuran-3-ylamino)methyl]phenyl}-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.60 (s, 1H), 8.15 (br s, 1H), 7.88 (br s, 1H), 7.65 (d, 1H), 7.40 (m, 3H), 7.26 (d, 1H), 6.96 (t, 1H), 6.90 (d, 1H), 4.60 (m, 3H), 3.90 (m, 2H), 3.01 (m, 3H); MS (EI) for C₂₁H₂₁N₅O₂: 376 (MH⁺).

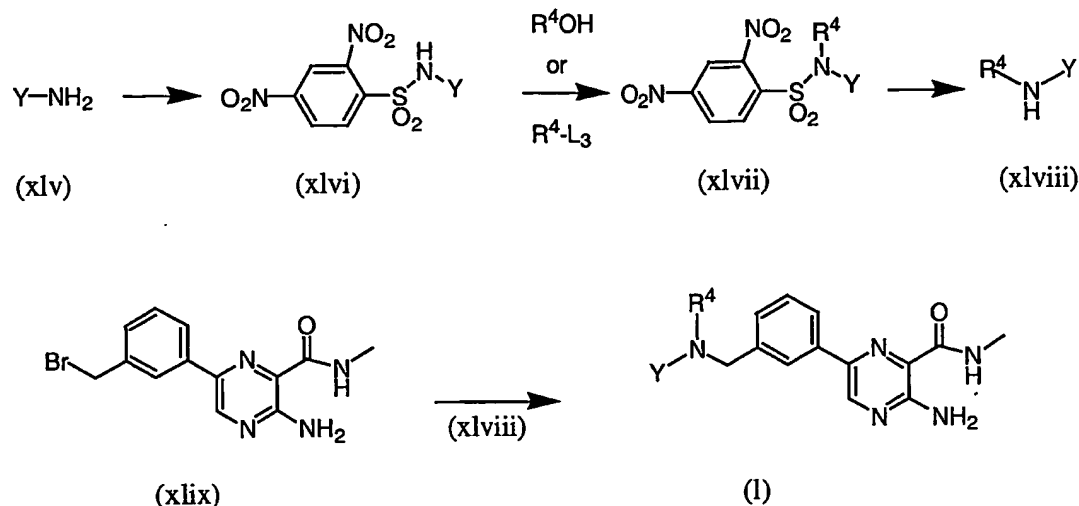
[0577] 3-amino-N-methyl-6-[3-([5-(4-methylphenyl)-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]pyrazine-2-carboxamide: ¹NMR (400 MHz, d₆-DMSO): 9.80 (br s, 1H), 9.02 (d, 1H), 8.82 (s, 1H), 8.60 (s, 1H), 8.21 (d, 1H), 7.50 (m, 7H), 7.25 (d, 1H), 4.82 (m, 1H), 4.30 (m, 2H), 3.20 (m, 1H), 2.92 (m, 1H), 2.80 (d, 3H), 2.50 (m, 1H), 2.30 (s, 3H), 2.40 (m, 1H); MS (EI) for C₂₉H₂₉N₅O: 464 (MH⁺).

Example 19

[0578] Scheme 10 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene, X is -CH₂NR⁴-, and Y is various groups. Note, as defined above any of W, X, or Y are optionally substituted; this is only a set of examples. As depicted, primary amine (xlv), which in this example incorporates Y, is incorporated into a compound of the invention, (I). For example, amine (xlv), which has Y as part of its structure for example, is converted to 2,4-dinitrosulfonamide (xlvi). Sulfonamide (xlvi) is used to make sulfonamide (xlvii). The transformation (xlvi) to (xlvii) is carried out either under alkylation (R⁴L₃, where -L₃ is a leaving group) or Mitsunobu (R⁴OH) conditions. The sulfonamide bond is cleaved, typically but not necessarily via addition of a primary amine e.g. *n*-propyl amine, to free secondary amine (xlviii), which has Y and R⁴ as part of its structure. Amine (xlviii) was then incorporated into compounds of the invention,

typically but not necessarily, via coupling to a benzyl bromide, for example (xlix), to make compound (I).

Scheme 10



3-amino-6-[3-({[(2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide

[0579] N-[(S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl]-2,4-

dinitrobenzenesulfonamide: To a solution of (S)-6-bromo-2-aminotetraline (0.20g, 0.9 mmol) in tetrahydrofuran (2 mL) was added N,N-diisopropylethylamine at 0° C. A solution of 2,4-dinitrobenzenesulfonyl chloride (0.29g, 1.1 mmol) in tetrahydrofuran (2 mL) was added slowly *via* a syringe. The reaction was allowed to warm to room temperature and further stirred until the completion of the reaction (30 min) as monitored by thin layer chromatography. The crude reaction mixture was poured into a separatory funnel and diluted with ethyl acetate (250 mL) and washed with water (30 mL), saturated aqueous sodium chloride (40 mL), and dried over anhydrous magnesium sulfate. Filtration and concentration afforded the crude product. A pure N-[(2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl]-2,4-dinitrobenzenesulfonamide (0.28g, 70% yield) was separated from trituration with ether/hexane (5:1, 30 mL) as a yellow solid. ¹NMR (400 MHz, CD₃OD): 8.75 (s, 1H), 8.60 (dd, 1H), 8.38 (d, 1H), 7.22 (br s, 1H), 7.20 (d, 1H), 6.90 (d, 1H), 3.65 (m, 1H), 2.65 (m, 4H), 1.90 (m, 1H), 1.75 (m, 1H).

[0580] 3-Amino-6-[3-({[(2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-

yl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: To a solution of N-[(2S)-6-

bromo-1,2,3,4-tetrahydronaphthalen-2-yl]-2,4-dinitrobenzenesulfonamide (68 mg, 0.15 mmol) in N,N-dimethylformamide (1mL) were added solid potassium carbonate (0.10 g, 0.75 mmol) and 3-amino-6-[3-(bromomethyl)phenyl]-N-methylpyrazine-2-carboxamide (70 mg, 0.22 mmol). The reaction stirred overnight at room temperature and diluted with ethyl acetate (50 mL). The organic layer was washed with water (30 mL) and saturated aqueous sodium chloride (30 mL), and dried over magnesium sulfate. Filtration and concentration gave crude product that was used in the next step without any purification.

[0581] To the dichloromethane solution (2 mL) of the above product was added excess n-propylamine (0.5mL) and stirred for 30 min. The organic solvent and excess reagent were removed at reduced pressure. Column purification on silica (9:1, ethyl acetate/methanol) gave an oily product.

[0582] To a solution of the above product in methanol (2 mL) was added 4.0 N hydrochloric acid (1,4-dioxane solution, 1.50 mL) and stirred for 40 min. Evaporation of the excess reagent and solvent, washing with diethyl ether afforded 3-amino-6-[3-(((2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide (16 mg, 23% yield) as hydrochloric acid salt. ¹NMR (400 MHz, CDCl₃): 8.62 (s, 1H), 8.02 (br s, 1H), 7.82 (br s, 1H), 7.75 (d, 1H), 7.40 (m, 2H), 7.20 (m, 2H), 6.74 (d, 1H), 4.01 (m, 2H), 2.90 (m, 6H), 2.60 (m, 2H), 2.12 (m, 1H), 1.70 (m, 1H); MS (EI) for C₂₃H₂₄N₅OBr: 466 (MH⁺).

[0583] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[0584] 3-amino-6-[3-(((2R)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.60 (s, 1H), 8.05 (br s, 1H), 7.83 (br s, 1H), 7.75 (d, 1H), 7.42 (m, 2H), 7.20 (m, 2H), 6.72 (d, 1H), 4.01 (m, 2H), 3.10 (m, 4H), 2.84 (m, 2H), 2.60 (m, 1H), 2.10 (m, 1H), 1.70 (m, 2H); MS (EI) for C₂₃H₂₄N₅OBr: 466 (MH⁺).

[0585] 3-amino-6-[3-[(cyclopentylamino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.70 (s, 1H), 8.48 (br s, 1H), 8.16 (d, 1H), 7.58 (m, 3H), 4.30 (s, 2H), 3.04 (br s, 3H), 2.20 (m, 2H), 1.72 (m, 7H); MS (EI) for C₁₈H₂₃N₅O: 326 (MH⁺).

[0586] 3-amino-N-methyl-6-{3-[[{(1S,2S)-2-[(phenylmethyl)oxy]cyclopentyl}amino)methyl]phenyl}pyrazine-2-carboxamide: ^1NMR (400 MHz, CDCl_3): 8.60 (s, 1H), 8.06 (br s, 1H), 7.80 (br s, 1H), 7.74 (d, 1H), 7.40 (t, 2H), 7.32 (m, 6H), 4.50 (q, 2H), 3.85 (m, 3H), 3.20 (m, 1H), 3.01 (d, 3H), 2.01 (m, 1H), 2.10 (m, 2H), 1.73 (m, 4H), 1.42 (m, 1H); MS (EI) for $\text{C}_{25}\text{H}_{29}\text{N}_5\text{O}_2$: 432 (MH^+).

[0587] 3-amino-N-methyl-6-{3-[[{(1R,2R)-2-[(phenylmethyl)oxy]cyclopentyl}amino)methyl]phenyl}pyrazine-2-carboxamide: ^1NMR (400 MHz, CDCl_3): 8.65 (s, 1H), 8.40 (br s, 1H), 8.12 (d, 1H), 7.58 (t, 1H), 7.48 (d, 1H), 7.34 (m, 5H), 4.60 (q, 2H), 4.32 (s, 2H), 4.28 (m, 1H), 3.60 (m, 1H), 3.02 (s, 3H), 2.30 (m, 1H), 2.15 (m, 1H), 1.83 (m, 4H); MS (EI) for $\text{C}_{25}\text{H}_{29}\text{N}_5\text{O}_2$: 432 (MH^+).

Example 20

3-Amino-6-[3-([2-(dimethylamino)-1-phenylethyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide

[0588] To a solution of (2-amino-2-phenethyl)dimethylamine (200 mg, 1.22 mmol) in acetonitrile (15 mL) were added 3-amino-6-[3-(bromomethyl)phenyl]-N-methylpyrazine-2-carboxamide (0.11 g, 0.35 mmol) and N,N-diisopropylethylamine (0.14 g, 1.05 mmol) at room temperature. The reaction was stirred overnight. Diluted with ethyl acetate (100 mL), washed with water (30 mL), saturated aqueous sodium chloride (brine), and dried over anhydrous magnesium sulfate. Filtration and concentration and reverse phase HPLC purification gave a trifluoro acetate salt. The fraction from the HPLC was neutralized with saturated aqueous sodium bicarbonate (50 mL) and diluted with ethyl acetate (200 mL). The organic layer was washed saturated aqueous sodium chloride (30 mL), dried over anhydrous magnesium sulfate, filtered, and concentrated to give the free amine. This free amine was converted into hydrochloric acid salt by treatment with 4.0 N hydrochloric acid (1,4-dioxane solution, 2.0 mL) in methanol solution (3 mL). Evaporation of the excess reagent and solvent gave 3-amino-6-[3-([2-(dimethylamino)-1-phenylethyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide (88 mg, 18% yield) as yellow solid: ^1NMR (400 MHz, CDCl_3): 8.61 (s, 1H), 8.05 (br s, 1H), 7.78 (br s, 1H), 7.74 (d, 1H), 7.35 (br m, 7H), 3.85 (d, 1H), 3.75 (dd, 1H), 3.60 (d, 1H), 3.04 (s, 3H), 2.65 (m, 2H), 2.20 (s, 6H); MS (EI) for $\text{C}_{23}\text{H}_{28}\text{N}_6\text{O}$: 405 (MH^+).

[0589] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[0590] 3-amino-N-methyl-6-(3-([(2-morpholin-4-yl-1-phenylethyl)amino]methyl}phenyl)pyrazine-2-carboxamide: ^1NMR (400 MHz, CDCl_3): 8.60 (s, 1H), 8.02 (br s, 1H), 7.74 (d, 1H), 7.70 (s, 1H), 7.39 (br m, 7H), 3.88 (d, 1H), 3.78 (dd, 1H), 3.65 (m, 3H), 3.58 (s, 1H), 3.05 (d, 3H), 2.55 (t, 1H), 2.45 (m, 2H), 2.30 (m, 3H); MS (EI) for $\text{C}_{25}\text{H}_{30}\text{N}_6\text{O}_2$: 447 (MH^+).

[0591] 3-amino-N-methyl-6-(3-([(1-phenyl-2-pyrrolidin-1-ylethyl)amino]methyl}phenyl)pyrazine-2-carboxamide: ^1NMR (400 MHz, CDCl_3): 8.60 (s, 1H), 8.12 (br s, 1H), 7.80 (s, 1H), 7.74 (d, 1H), 7.40 (br m, 5H), 7.30 (m, 2H), 3.82 (d, 1H), 3.74 (d, 1H), 3.60 (d, 1H), 3.05 (d, 3H), 2.90 (m, 2H), 2.70 (br s, 1H), 2.50 (m, 1H), 2.40 (m, 1H), 2.25 (dd, 2H), 1.70 (m, 4H); MS (EI) for $\text{C}_{25}\text{H}_{30}\text{N}_6\text{O}$: 431 (MH^+).

Example 21

3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-hydroxyethyl)pyrazine-2-carboxamide

[0592] To a solution of methyl 3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-hydroxyethyl)pyrazine-2-carboxylate (30 mg, 08 mmol) in methanol (1 mL) was added ethanolamine (20 mg, 0.32 mmol). The reaction was heated at 80-90° C overnight. The reaction cooled to room temperature and diluted with ethyl acetate (150 mL). The organic layer was washed with saturated aqueous sodium bicarbonate and saturated aqueous sodium chloride, and dried over magnesium sulfate. Filtration, concentration, and column purification on silica (9.5:0.5, dichloromethane/methanol) afforded free amine product. To a solution of the amine in diethyl ether (2 mL) was added 4 N hydrochloric acid (1,4-dioxane solution, 1 mL) and stirred for 30 min. Concentration and washings with diethyl ether afforded 3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-hydroxyethyl)pyrazine-2-carboxamide (13 mg, 38% yield) as yellow hydrochloric acid salt. ^1NMR (400 MHz, CDCl_3): 8.58 (s, 1H), 8.40 (br s, 1H), 7.90 (br s, 1H), 7.68 (m, 1H), 7.38 (m, 3H), 7.20 (m, 3H), 4.35 (t, 1H), 3.90 (d, 2H), 3.80 (t, 2H), 3.60 (m, 2H), 3.06 (m, 1H), 2.84 (m, 1H), 2.48 (m, 3H), 1.92 (m, 1H); MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_5\text{O}_2$: 404 (MH^+).

[0593] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[0594] 3-amino-N-cyclopropyl-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)pyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 8.60 (s, 1H), 8.05 (br s, 1H), 7.83 (br s, 1H), 7.70 (m, 1H), 7.40 (m, 3H), 7.20 (m, 3H), 4.35 (t, 1H), 4.02 (d, 2H), 3.02 (m, 1H), 2.85 (m, 2H), 2.50 (m, 1H), 1.92 (m, 1H), 1.13 (m, 1H), 0.90 (m, 2H), 0.70 (m, 1H); MS (EI) for C₂₄H₂₅N₅O: 400 (MH⁺).

[0595] 5-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine: ¹NMR (400 MHz, CDCl₃): 8.57 (s, 1H), 7.95 (br s, 1H), 7.78 (m, 1H), 7.42 (m, 3H), 7.24 (m, 3H), 4.35 (t, 1H), 4.02 (d, 4H), 3.70 (m, 2H), 3.05 (m, 1H), 2.85 (m, 1H), 2.48 (m, 1H), 1.94 (m, 5H); MS (EI) for C₂₅H₂₇N₅O: 414 (MH⁺).

[0596] 3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(3-hydroxypropyl)pyrazine-2-carboxamide: ¹NMR (400 MHz, CDCl₃): 9.56 (s, 1H), 8.65 (m, 1H), 8.38 (br s, 1H), 7.72 (d, 1H), 7.38 (t, 2H), 7.50 (m, 4H), 4.40 (t, 1H), 4.01 (q, 2H), 3.60 (m, 3H), 3.40 (m, 1H), 3.08 (m, 1H), 2.90 (m, 1H), 2.50 (m, 1H), 2.00 (m, 1H), 1.75 (m, 2H); MS (EI) for C₂₄H₂₇N₅O₂: 418 (MH⁺).

Example 22

3-amino-6-(3-((1S)-1-((1S)-2,3-dihydro-1H-inden-1-ylamino)ethyl}phenyl)-N-methylpyrazine-2-carboxamide

[0597] N-[(1S)-1-(3-bromophenyl)ethyl]-2,4-dinitrobenzenesulfonamide: To a stirred ice-cooled solution of (S)-(-)-3-bromo- α -methylbenzylamine (832 mg, 4.16 mmol, from Chiragene) in 15.6 mL of THF was added 1.33 g (4.99 mmol, 1.20 eq.) of 2,4-dinitrobenzenesulfonyl chloride, followed by 1.08 mL (6.20 mmol, 1.49 eq.) of N,N-diisopropylethylamine. The ice bath was removed, and the mixture was stirred at room temperature for 19 h and then concentrated. The residue was taken up in EtOAc, washed with 2 x H₂O, dried over Na₂SO₄, filtered, and concentrated. The residue was sonicated in ca. 5 mL of EtOAc until a precipitate was formed. The mixture was diluted to ca. 25 mL with hexanes. Filtration afforded product as a tan solid (1.69 g, 94.4% yield).

[0598] N-[(1S)-1-(3-bromophenyl)ethyl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-2,4-dinitrobenzenesulfonamide: A flask was charged with 500 mg (1.16 mmol) of N-[(1S)-1-

(3-bromophenyl)ethyl]-2,4-dinitrobenzenesulfonamide, 625 mg (4.66 mmol, 4.02 eq.) of (R)-(-)-1-indanol (from Sigma-Aldrich), 305 mg (1.16 mmol, 1.0 eq.) of triphenylphosphine, 6.75 mL of benzene and the mixture cooled on an ice-water bath. To the mixture was added 0.183 mL (1.16 mmol, 1.0 eq.) of diethyl azodicarboxylate (DEAD). The ice bath was removed, and stirring was continued for 1 h. The mixture was cooled on an ice-water bath, followed by addition of 305 mg (1.16 mmol, 1.0 eq.) of triphenylphosphine and 0.183 mL (1.16 mmol, 1.0 eq.) of DEAD. The ice bath was removed, and stirring continued for 1 h. The mixture was cooled on an ice-water bath, followed by a third addition of 305 mg (1.16 mmol, 1.0 eq.) of triphenylphosphine and 0.183 mL (1.16 mmol, 1.0 eq.) of DEAD. The ice bath was removed, and stirring continued for 1 h. The mixture was diluted with EtOAc, washed with 2 x sat. aqueous NaCl, dried (Na_2SO_4), filtered, and concentrated. The crude material was purified by silica gel chromatography on a 30 mm column (3.5 inches of silica) using EtOAc as eluent. The product was triturated in 10% EtOAc in hexanes and filtered affording 625 mg (98.5% yield) of product as a solid.

[0599] [(1S)-1-(3-bromophenyl)ethyl](1S)-2,3-dihydro-1H-inden-1-ylamine: To a mixture of 625 mg (1.14 mmol) of N-[(1S)-1-(3-bromophenyl)ethyl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-2,4-dinitrobenzenesulfonamide in 11.5 mL of CH_2Cl_2 was added 1.44 mL (17.5 mmol, 15.4 eq.) of n-propylamine. The reaction was stirred for 16 h at room temperature and concentrated. The crude material was purified by silica gel chromatography on a 25 mm column (3.5 inches of silica gel) using 10% EtOAc in hexanes as eluent. After concentration, 4.0 mL of Et_2O was added to the residue, followed by 4 N HCl in dioxane. The mixture was sonicated, filtered, and washed with Et_2O . The material was taken up in EtOAc, washed with 2 x sat. aqueous NaHCO_3 , dried (Na_2SO_4), filtered, and concentrated to afford 170 mg (47.2% yield) of product.

[0600] (1S)-N-[(1S)-1-[3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)ethyl]indane-1-amine was synthesized according to the procedure described in above from [(1S)-1-(3-bromophenyl)ethyl](1S)-2,3-dihydro-1H-indene-1-ylamine and bis(pinacolato)diboron. Yield 64%. ^1H NMR (400 MHz, CDCl_3): 7.80 (br s, 1H), 7.72 (br d, 1H), 7.58 (br d, 1H), 7.35 (m, 2H), 7.18 (m, 3H), 4.12 (m, 2H), 2.90 (m, 1H), 2.70 (m, 1H), 2.20 (m, 1H), 1.70 (m, 1H), 1.40 (d, 3H), 1.24 (s, 12H).

[0601] 3-amino-6-(3-[(1S)-1-[(1S)-2,3-dihydro-1H-inden-1-ylamino]ethyl]phenyl)-N-methylpyrazine-2-carboxamide was prepared according to the procedure described above

by coupling (1S)-N-[(1S)-1-[3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan-2-yl)ethyl]indane-1-amine and 3-amino-6-bromo-N-methyl-2-pyrazincarboxamide as hydrochloric acid salt. Yield 34 %. ¹H NMR (400 MHz, CDCl₃): 8.64 (s, 1H), 8.15 (br s, 1H), 8.02 (br s, 1H), 7.78 (m, 1H), 7.49 (m, 2H), 7.40 (m, 1H), 7.20 (m, 4H), 4.21 (m, 2H), 3.02 (m, 4H), 2.78 (m, 1H), 2.22 (m, 1H), 1.80 (br s, 1H), 1.42 (d, 3H); MS (EI) for C₂₃H₂₅N₅O: 388 (MH⁺).

Example 23

3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl)phenylpyrazine-2-carboxamide

[0602] 3-bromo-N-[(1S)-2,3-dihydro-1H-indene-1-yl]benzenesulfonamide was synthesized according to the procedure described above from (S)-1-aminoindan and 3-bromobenzenesulfonyl chloride. Yield 52%. ¹H NMR (400 MHz, CDCl₃): 7.20 (m, 1H), 7.90 (dd, 1H), 7.80 (dd, 1H), 7.50 (t, 1H), 7.20 (m, 2H), 7.15 (m, 1H), 7.02 (br d, 1H), 4.80 (t, 1H), 2.90 (m, 1H), 2.70 (m, 1H), 2.20 (m, 1H), 1.70 (m, 1H).

[0603] N-[(1S)-2,3-dihydro-1H-inden-1-yl]-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan)benzenesulfonamide was synthesized according to the procedure described above from 3-Bromo-N-[(1S)-2,3-dihydro-1H-indene-1-yl]benzenesulfonamide and bis(pinacolato)diboron. Yield 58 %. ¹H NMR (400 MHz, CDCl₃): 8.40 (br s, 1H), 8.02 (m, 2H), 7.54 (t, 1H), 7.16 (m, 4H), 4.90 (m, 1H), 4.70 (d, 1H), 2.90 (m, 1H), 2.78 (m, 1H), 2.30 (m, 1H), 1.70 (m, 1H), 1.24 (s, 12H).

[0604] 3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl)phenylpyrazine-2-carboxamide was prepared according to the procedure described above by coupling N-[(1S)-2,3-dihydro-1H-inden-1-yl]-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan)benzenesulfonamide and 3-amino-6-bromo-N-methyl-2-pyrazincarboxamide. Yield 50%. ¹H NMR (400 MHz, d₆-DMSO): 8.86 (s, 1H), 8.60 (br s, 1H), 8.45 (br d, 1H), 8.25 (br s, 1H), 8.20 (d, 1H), 7.85 (t, 2H), 7.70 (t, 1H), 7.19 (m, 4H), 4.80 (m, 1H), 2.80 (m, 1H), 2.62 (m, 1H), 2.00 (m, 1H), 1.60 (m, 1H); MS (EI) for C₂₀H₁₉N₅O₃S: 410 (MH⁺).

[0605] 3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl)phenyl-N-piperidin-3-ylpyrazine-2-carboxamide was prepared according to the procedure described above by coupling N-[(1S)-2,3-dihydro-1H-inden-1-yl]-3-(4,4,5,5-tetramethyl-1,3,2-dioxaborolan)benzenesulfonamide and tert-butyl 3-[2-(3-amino-6-bromopyrazin-2-yl)-

2-oxoethyl]piperidine-1-carboxylate followed by the deprotection of the boc group with 4.0 N hydrochloric acid(1,4-dioxane solution). Yield 50 %. ¹H NMR (400 MHz, CD₃OD): 8.78 (br s, 1H), 8.58 (br s, 1H), 8.35 (d, 1H), 7.95 (d, 1H), 7.75 (t, 1H), 7.18 (m, 2H), 7.10 (m, 2H), 4.85 (t, 1H), 4.30 (br s, 1H), 3.72 (m, 1H), 3.68 (m, 1H), 3.55 (m, 1H), 3.50 (br d, 1H), 3.35 (dr d, 1H), 3.15 (br t, 1H), 3.00 (br t, 1H), 2.90 (m, 1H), 2.70 (m, 1H), 2.10 (m, 2H), 1.78 (m, 1H), 1.70 (m, 1H); MS (EI) for C₂₅H₂₈N₆O₃S: 493 (MH⁺).

[0606] N-[3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)phenyl]-2-(2,4-dichlorophenyl)acetamide: ¹H NMR (400 MHz, CD₃OD): 8.52 (br s, 1H), 8.48 (br s, 1H), 7.82 (br d, 1H), 7.48 (m, 6H), 7.35 (m, 1H), 3.92 (s, 3H), 3.70 (m, 1H), 3.42 (m, 2H), 3.05 (m, 1H), 2.30 (m, 1H), 2.02 (m, 3H); MS (EI) for C₂₅H₂₄N₈OCl₂: 524 (MH⁺).

[0607] 1,1-dimethylethyl (3R)-3-[[3-(5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)phenyl]carbonyl]amino}pyrrolidine-1-carboxylate: ¹H NMR (400 MHz; DMSO-D₆): 8.91 (s, 1H); 8.86 (m, 1H); 8.63 (d 1H); 8.45 (s, 1H); 8.38 (d, 1H); 7.82 (d, 1H); 7.76 (t, 1H); 4.45 (m, 1H); 3.7-3.4 (m, 4H); 2.85 (d, 3H); 2.33 (m, 1H); 1.9 (m, 1H); 1.4 (s, 9H); MS (EI) for C₂₂H₂₈N₆O₄: 441 (MH⁺)

Example 24

3-amino-N-methyl-6-(3-[[3-(3R)-pyrrolidin-3-ylamino]carbonyl]phenyl)pyrazine-2-carboxamide

[0608] 1,1-dimethylethyl-3-[[3-(5-amino-6[(methylamino)carbonyl]pyrazin-2-yl)phenyl]carbonyl]amino}pyrrolidine-1-carboxylate (0.7 g, mmol) was suspended in dichloromethane (4 mL) and TFA (3 mL) was added. The mixture was stirred at ambient temperature for 2 hr. After removal of volatiles under reduced pressure, the residue was partitioned between saturated solution K₂CO₃ (300 mg) and dichloromethane (10 mL). The organic layer was separated, dried (Na₂SO₄) and evaporated to dryness to afford 0.4 g (74 % yield) of 3-amino-N-methyl-6-(3-[[3-(3R)-pyrrolidin-3-ylamino]carbonyl]phenyl)pyrazine-2-carboxamide. ¹H NMR (400 MHz; DMSO-D₆): 8.9 (s, 1H); 8.85 (m, 1H); 8.71 (d, 1H); 8.48 (m, 1H); 7.4 (d, 1H); 7.84 (d, 1H); 7.6 (t, 1H); 4.54 (m, 2H); 3.6-3.18 (m, 4H); 2.85 (d, 3H); 2.25 (m, 1H); 2.1 (m, 1H); MS (EI) for C₁₇H₂₀N₆O₂: 341 (MH⁺)

[0609] 3-{[(4-chlorophenyl)methyl]amino}-6-{3-[[[(3R)-1-[(4-chlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: To a solution of 3-amino-N-methyl-6-(3-[[[(3R)-pyrrolidin-3-ylamino]carbonyl]phenyl]pyrazine-2-carboxamide (34.0mg, 0.1 mmol) in acetonitrile was added K₂CO₃ (27.6 mg, 0.2 mmol) followed by 4-chlorobenzyl bromide (30.7 mg, 0.15 mmol). The mixture was allowed to stir at room temperature over 12 h and filtered. The acetonitrile solution was evaporated to dryness, the residue was chromatographed on silica gel using ethyl acetate/ hexane 4:1 v/v and trituration of the resulting solid with hexane afforded 3-{[(4-chlorophenyl)methyl]amino}-6-{3-[[[(3R)-1-[(4-chlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide (28 mg, 47.6 %yield): ¹H NMR (400 MHz; DMSO-D₆): 8.9 (s, 1H); 8.85 (m, 1H); 8.75 (m, 1H); 8.38 (m, 2H); 7.78-7.5 (m, 11H); 4.63 (m, 5H); 3.9 (m, 1H); 3.6 (m, 2H); 3.4 (m, 1H); 2.85 (d, 3H); 2.5 (m, 1H); 2.2 (m, 1H); MS (EI) for C₃₁H₃₀N₆O₂Cl₂: 589 (MH⁺)

[0610] The following compounds were prepared in an analogous manner as 3-{[(4-chlorophenyl)methyl]amino}-6-{3-[[[(3R)-1-[(4-chlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide.

[0611] 3-amino-6-{3-[[[(3R)-1-[(2,4-dichlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz; DMSO-D₆): 8.92 (s, 1H); 8.86 (m, 1H); 8.6 (d, 1H); 8.46 (m, 1H); 8.36 (d, 1H); 7.82 (d, 1H); 7.6-7.4 (m, 4H); 4.43 (m, 1H); 3.7 (m, 2H); 2.9 (m, 1H); 2.88 (d, 3H); 2.75 (m, 1H); 2.58 (m, 2H); 2.2 (m, 1H); 1.85 (m, 1H); MS (EI) for C₂₄H₂₄N₆O₂Cl₂: 499 (MH⁺)

[0612] N-methyl-3-([3-(methyloxy)phenyl]methyl)amino)-6-(3-[[[(3R)-1-([3-(methyloxy)phenyl]methyl]pyrrolidin-3-yl)amino]carbonyl]phenyl)pyrazine-2-carboxamide: 8.98 (m, 1H); 8.92 (m, 1H); 8.88 (s, 1H); 8.48 (m, 1H); 8.38 (d, 1H); 7.78 (d, 1H); 7.6-7.0 (m, 9H); 4.7 (m, 5H); 3.95 (m, 1H); 3.85 (s, 3H); 3.74 (s, 3H); 3.7 (m, 2H); 3.54 (m, 1H); 2.85 (s, 3H); 2.45 (m, 1H); 2.28 (m, 1H); MS (EI) for C₃₃H₃₆N₆O₄: 581 (MH⁺)

[0613] 3-Amino-6-{3-[[[(3R)-1-[(2,6-dichlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz; DMSO-D₆): 8.91 (s, 1H); 8.86 (m, 1H); 8.55 (d, 1H); 8.46 (m, 1H); 8.38 (d, 1H); 7.82 (d, 1H); 7.6-7.3 (m, 4H); 4.4 (m, 1H); 3.88 (s, 2H); 3.0 (m, 1H); 2.85 (d, 3H); 2.8 – 2.54 (m, 3H); 2.15 (m, 1H); 1.85 (m, 1H); MS (EI) for C₂₄H₂₄N₆O₂Cl₂: 499 (MH⁺)

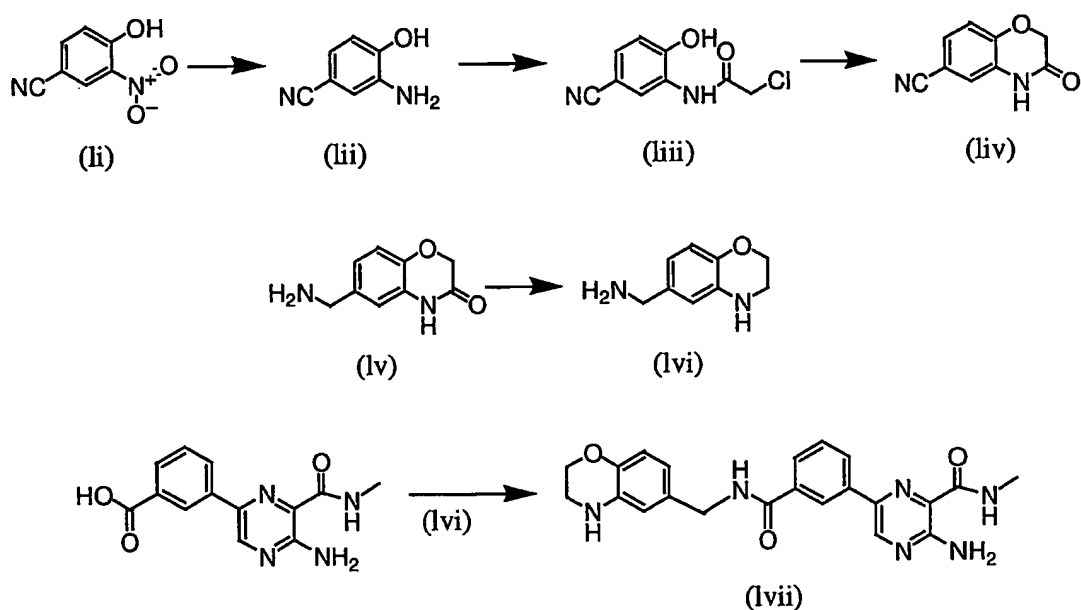
- [0614] N-methyl-3-[(phenylmethyl)amino]-6-[3-({[(3R)-1-(phenylmethyl)pyrrolidin-3-yl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.92 (m, 2H); 8.86 (s, 1H); 8.45 (s, 1H); 8.36 (d, 1H); 7.76 (d, 1H); 7.7-7.46 (m, 11H); 4.65 (m, 5H); 3.91 (m, 1H); 3.62 (m, 2H); 3.45 (m, 1H); 2.85 (d, 3H); 2.49 (m, 1H); 1.22 (m, 1H); MS (EI) for $\text{C}_{31}\text{H}_{32}\text{N}_6\text{O}_2$: 521 (MH^+)
- [0615] 1,1-dimethylethyl 3-({[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbonyl]amino}methyl)pyrrolidine-1-carboxylate: ^1H NMR (400 MHz; DMSO- D_6): 8.91 (s, 1H); 8.85 (m, 1H); 8.68 (m, 1H); 8.46 (s, 1H); 8.38 (d, 1H); 7.82 (d, 1H); 7.56 (t, 1H); 3.4 (m, 5H); 3.0 (m, 1H); 2.85 (d, 3H); 2.47 (m, 1H); 1.95 (m, 1H); 1.6 (m, 1H); 1.39 (s, 9H); MS (EI) for $\text{C}_{23}\text{H}_{30}\text{N}_6\text{O}_4$: 455 (MH^+)
- [0616] 1,1-dimethylethyl 2-({[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbonyl]amino}methyl)pyrrolidine-1-carboxylate: ^1H NMR (400 MHz; DMSO- D_6): 8.91 (s, 1H); 8.85 (m, 1H); 8.5 (m, 1H); 8.46 (s, 1H); 8.38 (d, 1H); 7.72 (d, 1H); 7.58 (t, 1H); 3.98 (m, 1H); 3.4 (m, 4H); 2.85 (d, 3H); 1.85 (m, 4H); 1.39 (s, 9H); MS (EI) for $\text{C}_{23}\text{H}_{30}\text{N}_6\text{O}_4$: 455 (MH^+)
- [0617] 3-amino-N-methyl-6-(3-({[(pyrrolidin-3-ylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.91 (s, 1H); 8.85 (m, 1H); 8.75 (m, 1H); 8.66 (br s, 2H); 8.48 (s, 1H); 8.38 (d, 1H); 7.84 (d, 1H); 7.58 (t, 1H); 3.6-3.2 (m, 5H); 3.1 (m, 1H); 2.91 (m, 1H); 2.85 (d, 3H); 2.6 (m, 1H); 2.0 (m, 1H); 1.7 (m, 1H); MS (EI) for $\text{C}_{18}\text{H}_{22}\text{N}_6\text{O}_2$: 355 (MH^+)
- [0618] 3-amino-N-methyl-6-(3-({[(pyrrolidin-2-ylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.91 (s, 1H); 8.85 (m, 1H); 8.5 (s, 1H); 8.4 (d, 1H); 7.84 (d, 1H); 7.58 (t, 1H); 3.6-3.4 (m, 3H); 3.1-3.2 (m, 2H); 2.85 (d, 3H); 2.0-1.6 (m, 4H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_6\text{O}_2$: 355 (MH^+)

Example 25

- [0619] Scheme 11 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene, X is -C(=O)NHCH₂-, and Y is a 3,4-dihydro-2H-benzo[1,4]oxazinyl group. As depicted, intermediate (li) is incorporated into a compound of the invention, (lvii). Nitro phenol (li) was reduced to aniline (lii). Then the aniline nitrogen was acylated to give intermediate

(liii). A ring closure was effected to yield intermediate (liv). Reduction of the cyano group gave (lv), followed by reduction of the carbonyl giving primary amine (lvi). Amine (lvi) was coupled to an acid to yield compounds of the invention, for example (lvii) as depicted below.

Scheme 11



3-amino-6-(3-[[[(3,4-dihydro-2H-1,4-benzoxazin-6-yl)methyl]amino]carbonyl]phenyl)-N-methyl pyrazine-2-carboxamide

[0620] 3-Amino-4-hydroxybenzonitrile: To a solution of 3-nitro-4-hydroxybenzonitrile (3.28 g, 20 mmol) in ethyl acetate was hydrogenated over 5% Pd/C, at an initial pressure 20 psi in a Parr shaker. After 3 h the reaction mixture was filtered (Celite). The solvent was removed under reduced pressure and the residue oil 3-Amino-4-hydroxybenzonitrile (2.6 g, 97% yield) used in the next step without further purification: ^1H NMR (400 MHz; DMSO- D_6): 6.85 (m, 2H); 6.73 (d, 1H); 5.0 (br s, NH).

[0621] 3-Oxo-3,4-dihydro-2H-1,4-benzoxazin-6-carbonitrile: 3-amino-4-hydroxybenzonitrile (g, 10 mmol) was dissolved in acetone (20 mL) and water (20 mL) containing sodium bicarbonate (4 g). 2-chloroacetyl chloride was added slowly and the mixture heated to reflux for 4 h and then allowed to stir overnight at 25 $^\circ\text{C}$. The layers were separated, and the water layer was extracted with ethyl acetate. The organic layers were combined and evaporated to give 3-oxo-3,4-dihydro-2H-1,4-benzoxazin-6-

carbonitrile as an oil (1.3 g): ^1H NMR (400 MHz; DMSO- D_6): 7.4 (d, 1H); 7.2 (s, 1H); 7.1 (d, 1H).

[0622] 3,4-Dihydro-2H-1,4-benzoxazine-6-ylmethyl)amine: a solution of 3-oxo-3,4-dihydro-2H-1,4-benzoxazin-6-carbonitrile (1.3 g) in anhydrous THF (20 mL) was slowly added to a stirred and ice-cooled solution of lithium aluminum hydride (30 mL, 1 M solution in THF). With continued cooling water (5 mL) and sodium hydroxide (5 mL, 20% solution), water (20 mL) and ethyl acetate (30 mL) were added in succession. The organic solution was decanted from the white residue. The residue was washed twice with ether, all the organic solution were combined then dried over anhydrous sodium sulfate. Filtration and concentration followed by drying in vacuo afforded 3,4-dihydro-2H-1,4-benzoxazine-6-ylmethylamine (0.78 g): ^1H NMR (400 MHz; DMSO- D_6): 6.52 (d, 1H); 6.49 (s, 1H); 6.37 (d, 1H); 5.6 (br s, NH); 4.05 (t, 2H); 3.4 (m, 2H).

[0623] 3-amino-6-(3-([(3,4-dihydro-2H-1,4-benzoxazin-6-ylmethyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz; DMSO- D_6): 8.96 (t, 1H); 8.79 (s, 1H); 8.81 (m, 1H); 8.5 (s, 1H); 8.35 (d, 1H); 7.85 (d, 1H); 7.65 (br s, 2H); 7.55 (t, 1H); 6.55 (m, 2H); 6.44 (d, 1H); 5.77 (s, 1H); 4.4 (d, 2H); 4.15 (m, 2H); 3.2 (s, 1H); 2.85 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{22}\text{N}_6\text{O}_3$: 419 (MH^+)

Example 26

3-amino-N-methyl-6-{3-([(1-(phenylmethyl)pyrrolidin-2-yl)methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide

[0624] To a stirred solution of 3-amino-N-methyl-6-(3-([(pyrrolidin-2-ylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide (35.4 mg, 0.01 mmol) and benzaldehyde (24.8 mg, 0.02 mmol) in methanol (3 mL) was added sodium cyanoborohydride (25 mg, 0.04 mmol). The mixture was stirred for 1 h and then acetic acid added until the pH of the solution was 7.0. The solution was stirred for an additional 6 h, solvent was removed and residue was partitioned with CHCl_3 and 2 M aqueous sodium hydroxide (5 mL). The aqueous phase was extracted with additional CHCl_3 (2x5 mL) and the combined organic layer were washed with saturated aqueous sodium chloride then dried over anhydrous sodium sulfate. Filtration, concentration and column chromatography provided 10mg of 3-amino-N-methyl-6-{3-([(1-(phenylmethyl)pyrrolidin-2-yl)methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide:

^1H NMR (400 MHz; CDCl_3): 8.7 (s, 1H); 8.65 (s, 1H); 8.28 (br s, 1H); 7.95 (d, 1H); 7.22 (d, 1H); 7.53 (t, 1H); 7.2-7.4 (m, 5H); 4.3 - 3.5 (m, 7H); 2.95 (d, 3H); 2.35 - 1.8 (m, 4H); MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2$: 445 (MH^+)

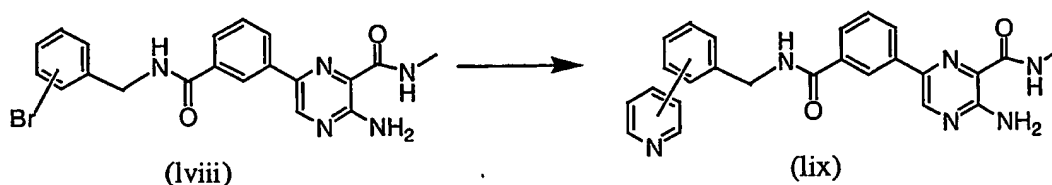
[0625] 3-amino-N-methyl-6-[3-([1-(phenylmethyl)pyrrolidin-3-yl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; CDCl_3): 8.67 (s, 1H); 8.42 (s, 1H); 8.15 (m, 1H); 7.95 (d, 1H); 7.84 (d, 1H); 7.53 (t, 1H); 7.4 (m, 5H); 4.15 (m, 2H); 3.75 (m, 1H); 3.4 (m, 2H); 3.0 (m, 5H); 2.22 (m, 1H); 1.95 (m, 1H); MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2$: 445 (MH^+)

[0626] 3-amino-N-methyl-6-[3-([(2-pyridin-4-ylphenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; $\text{DMSO}-d_6$): 9.1 (t, 1H); 8.6 (s, 1H); 8.84 (m, 1H); 8.64 (m, 1H); 8.42 (m, 1H); 7.35 (d, 1H); 7.8 (d, 1H); 7.6 - 7.36 (m, 6H); 7.28 (d, 1H) 4.43 (d, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{25}\text{H}_{22}\text{N}_6\text{O}_2$: 439 (MH^+)

Example 27

[0627] Scheme 12 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene, X is -C(=O)NHCH₂-, and Y is a bis-aryl, aryl-heteroaryl, or heteroaryl-heteroaryl group. As depicted, intermediate (lviii), a compound of the invention, is converted to (lix), another compound of the invention, for example, via aromatic coupling reactions known in the art.

Scheme 12



[0628] 3-amino-N-methyl-6-[3-([(2-pyridin-3-ylphenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz; $\text{DMSO}-d_6$): 9.1 (t, 1H); 8.6 (s, 1H); 8.64 (m, 1H); 8.58 (m, 1H); 8.42 (s, 1H); 8.34 (d, 1H); 7.88 (m, 1H); 7.8 (d, 1H); 7.58 - 7.34 (m, 5H); 7.28 (d, 1H) 4.43 (d, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{25}\text{H}_{22}\text{N}_6\text{O}_2$: 439 (MH^+)

[0629] 3-amino-N-methyl-6-[3-([(4-pyridin-3-ylphenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: (400 MHz; CD₃OD): 8.24 (t, 1H); 9.08 (m, 1H); 8.78 (s, 1H); 8.72 (m, 2H); 8.55 (m, 1H); 8.2 (d, 1H); 8.0 (m, 1H); 7.88 (d, 1H); 7.76 (m, 2H); 7.58 (m, 3H); 4.54 (d, 2H); 2.85 (s, 3H); MS (EI) for C₂₅H₂₂N₆O₂: 439 (MH⁺)

Example 28

3-amino-6-[3-(1H-benzimidazol-2-yl)phenyl]-N-methylpyrazine-2-carboxamide

[0630] 3-Amino-6-(3-formylphenyl)-N-methylpyrazine-2-carboxamide: Methyl 3-amino-6-(3-formylphenyl)pyrazine-2-carboxylate (0.6 g, mmol) was suspended in ammonia (30 mL, 2.0 M solution in methyl alcohol) and brought to reflux over 2 h, then water (100 mL) was added. The solid was collected, treated with 4 M HCl in 1,4-dioxane (10 mL) and stirred at 70 °C overnight. The solvent was removed and the residue was partitioned with ethyl acetate and saturated sodium bicarbonate. The organic layer was washed with brine then dried over sodium sulfate. Filtration, concentration and column chromatography on silica afforded 3-amino-6-(3-formylphenyl)-N-methylpyrazine-2-carboxamide (0.4g).

[0631] 3-amino-6-[3-(1H-benzimidazol-2-yl)phenyl]-N-methylpyrazine-2-carboxamide: A solution of 3-amino-6-(3-formylphenyl)-N-methylpyrazine-2-carboxamide (25.4 mg, 0.1 mmol) and 1,2-phenyldiamine in NMP (3 mL) was heated at 80 °C for 8 h. The solvent was removed and the residue was triturated with 0.1 M aqueous hydrochloric acid. Filtration and hplc purification of the precipitate afforded (24 mg, 57% yield) of product: (400 MHz; DMSO-D₆): 8.9 (s, 1H); 8.85 (m, 1H); 8.8 (s, 1H); 8.31 (d, 1H); 8.2 (d, 1H); 7.6 (m, 3H); 7.2 (m, 2H); 2.85 (d, 3H); MS (EI) for C₁₉H₁₆N₆O: 345 (MH⁺)

Example 29

3-amino-N-methyl-6-[3-(4-phenyl-1H-imidazol-2-yl)phenyl]pyrazine-2-carboxamide

[0632] 3-amino-6-[3-[amino(hydroxyimino)methyl]phenyl]pyrazine-2-carboxamide: 3-Amino-6(3-cyanophenyl)-N-methylpyrazine-2-carboxamide (0.7 g, 2.76 mmol) and hydroxylamine (5 mL, 50% aqueous) were stirred in methanol/ THF (5 mL, 1 :1) at 80 °C for 8 h. The mixture was poured into water (30 mL). The solid was collected by filtration,

washed with ethyl ether and dried to give 3-amino-6-{3-[amino(hydroxyimino)methyl]phenyl}pyrazine-2-carboxamide (0.75 g, 94.9% yield) as a white solid: ^1H NMR (400 MHz; DMSO- d_6): 9.7 (s, 1H); 8.85 (m, 2H); 8.3 (s, 1H); 8.2 (d, 1H); 7.74 (d, 1H); 7.4 (t, 1H); 6.0 (s, 2H); 2.85 (d, 3H); MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_6\text{O}_2$: 287 (MH^+).

[0633] 3-amino-6-{3-[amino(imino)methyl]phenyl}-N-methylpyrazine-2-carboxamide:

To a stirred solution of 3-amino-6-{3-[amino(hydroxyimino)methyl]phenyl}pyrazine-2-carboxamide (0.858 mg, 3 mmol) in pyridine (5 mL) was added acetic anhydride (4 mmol). The mixture was poured into water (50 mL) and extracted with ethyl acetate (30 mL). The extract was washed with brine, dried (Na_2CO_3), and concentrate to provide 6-{3-[(E,Z)-[(acetyloxy)imino](amino)methyl]phenyl}-3-amino-N-methylpyrazine-2-carboxamide: To a solution of this crude material (400 mg) in 950 mL of methanol was added Pd/C (5%, 20 mg) and the mixture was hydrogenated at 30 psi on a Parr shaker. The mixture was filtered through celite and the filtrate was evaporated, and concentrated to provide (300 mg, 37% yield) of 3-amino-6-{3-[amino(imino)methyl]phenyl}-N-methylpyrazine-2-carboxamide: MS (EI) for $\text{C}_{13}\text{H}_{14}\text{N}_6\text{O}$: 271 (MH^+).

[0634] 3-amino-N-methyl-6-[3-(4-phenyl-1H-imidazol-2-yl)phenyl]pyrazine-2-

carboxamide: To a stirred solution of 3-amino-6-{3-[amino(imino)methyl]phenyl}-N-methylpyrazine-2-carboxamide (0.136 mg, 0.5 mmol) in DMF (5 mL) was added bromoacetophenone (0.11 g, 0.55 mmol) followed by K_2CO_3 (0.276 g). The resulting mixture was heated at 70 $^\circ\text{C}$ for 0.5 h. The mixture was poured into water (30 mL) and extracted with ethyl acetate (30 mL). The extract was washed with brine, dried (Na_2SO_4), and concentrated. The product was purified by hplc to provide 3-amino-N-methyl-6-[3-(4-phenyl-1H-imidazol-2-yl)phenyl]pyrazine-2-carboxamide (58 mg): ^1H NMR (400 MHz; DMSO- d_6): 8.94 (s, 1H); 8.82 (m, 1H); 8.62 (s, 1H); 8.2 (d, 1H); 8.04 (d, 1H); 7.9 – 7.78 (m, 3H); 7.58 (t, 1H); 7.38 (m, 2H); 7.24 (t, 1H); 2.82 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{18}\text{N}_6\text{O}$: 371 (MH^+).

Example 30

3-amino-N-methyl-6-[3-({[(4-propylphenyl)methyl]amino}methyl)phenyl]pyrazine-2-carboxamide

[0635] 3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)methyl]phenyl]

pyrazine-2-carboxamide: To a stirred solution of 3-amino-6-(3-formylphenyl)-N-methylpyrazine-2-carboxamide (50 mg, 0.19 mmol) and 4-propylbenzylamine (35.7 mg, 0.24 mmol) in THF (5 mL) was added sodium cyanoborohydride (ca. 5 eq.) and the solution was stirred until refluxed for 1 h. The solvent was removed under reduced pressure. The residue was partitioned between ethyl acetate (50 mL) and saturated aqueous solution of Na₂CO₃ (10 mL). The phases were separated. The organic layer was washed with saturated aqueous sodium chloride then dried (Na₂SO₄). Filtration and concentration followed by HPLC purification gave (24 mg, 32% yield) of 3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide:

¹H NMR (400 MHz; DMSO-d₆): 9.7 (br s, 1H); 9.0 (m, 1H); 8.87 (s, 1H); 8.5 (s, 1H); 8.4 (d, 1H); 7.5 (m, 6H); 7.43 (m, 2H); 4.2 (d, 2H); 2.82 (d, 3H); 2.56 (m, 2H); 1.6 (m, 2H); 0.9 (m, 3H); MS (EI) for C₂₃H₂₇N₅O: 390 (MH⁺).

[0636] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0637] 3-amino-6-[3-(((5-bromo-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-

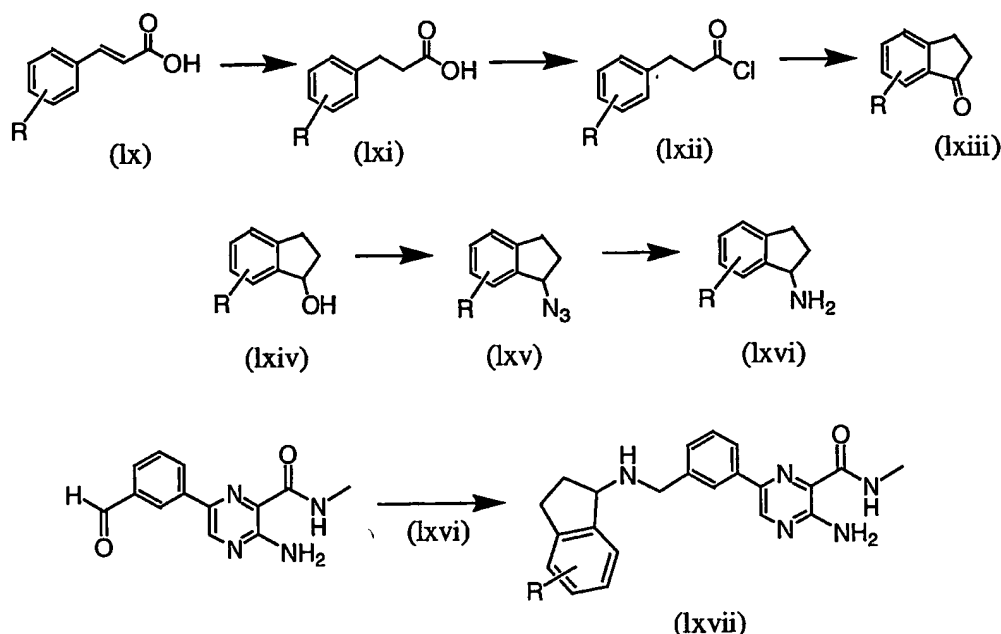
methylpyrazine-2-carboxamide: ¹H NMR (400 MHz; DMSO-d₆): 9.8 (br s, 1H); 9.0 (m, 1H); 8.88 (s, 1H); 8.62 (s, 1H); 8.21 (m, 1H); 8.01 (m, 3H); 7.68 (m, 1H); 7.5 (m, 2H); 7.3 (t, 1H); 4.2 (d, 2H); 2.85 (t, 3H); MS (EI) for C₂₀H₁₉N₅OBr: 446 (MH⁺).

Example 31

[0638] Scheme 13 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene, X is -C(=O)NHCH₂-, and Y is an indanyl group. As depicted, intermediate (Ix), is incorporated into (Ixxvii), a compound of the invention. 1-Amino-2,3-dihydro-R₁,R₂-substituted-1H-indene(s) were obtained from commercial sources or were prepared from cinnamic acids (or aryl propionic acids) by methods known to those skilled in the art as depicted in Scheme 13. Cinnamic acids (Ix) were hydrogenated to afford acids (Ixi). Acids (Ixi) were converted to the corresponding acyl chlorides and subjected to electrophilic aromatic substitution conditions to afford ketones (Ixxiii). Ketones (Ixxiii) were reduced to the corresponding alcohols (Ixxiv), and the alcohols subsequently converted to the corresponding azides (Ixxv). Azides (Ixxv) were reduced to form the

corresponding 1-aminoindanes (lxvi). Note, azides (lxv) with chloro- or bromo-substitution on phenyl ring portion were reduced to the 1-aminoindanes (lxvi) using SnCl_2 in ethyl acetate, typically. Aminoindanes (lxvi) were incorporated into compounds of the invention, e.g. (lxvii), via reductive amination, for example, as depicted. As mentioned, amines were also incorporated via coupling with corresponding benzyl bromides to give, for example, compounds (lxvii).

Scheme 13



[0639] 3-amino-6-(3-((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide: To a stirred solution of 3-amino-6-(3-formylphenyl)-N-methylpyrazine-2-carboxamide (50 mg, 0.19 mmol) and 1-amino-4,7-difluoroindane (67.6 mg, 0.4 mmol) in THF (5 mL) was added sodium cyanoborohydride (62.8 mg, 1 mmol) and 1 drop of acetic acid. The solution was stirred overnight and solvent was removed under reduced pressure. The residue was partitioned between ethyl acetate (50 mL) and saturated aqueous solution of Na_2CO_3 (10 mL). The phases were separated. The organic layer was washed with HCl (5 mL 0.5 M) then saturated aqueous sodium chloride and dried (Na_2SO_4). Filtration and concentration follow by hplc purification gave (42.2 mg, 50% yield) of 3-amino-6-(3-((4,7-difluoro-3-methyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide. An excess of HCl in dioxane (0.2 mL, 4.0 M) was added to the solution and the solvent was removed to give hydrochloride: ^1H NMR (400 MHz; $\text{DMSO}-d_6$): δ 0.1 (br s, 1H); 9.7 (br s, 1H); 9.0 (m,

1H); 8.85 (s, 1H); 8.2 (m, 1H); 7.5 (m, 2H); 7.3 – 7.2 (m, 2H); 5.0 (m, 1H); 4.6 – 4.15 (m, 3H); 3.4 (m, 1H); 2.95 (m, 1H); 2.85 (d, 3H); 2.6 (m, 1H); 2.46 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OF_2$ HCl: 410 (MH^+).

[0640] 3-amino-6-(3-([(4,7-difluoro-3-methyl-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6) 9.0 (m, NH); 8.88 (s, 1H); 8.6 (s, 1H); 8.2 (s, 1H); 7.5 (m, 2H); 7.4 (m, 2H); 4.5 (m, 1H); 4.2 (m, 2H); 2.85 (m, 3H); 2.8 (m, 1H); 2.2 (m, 2H); 1.3 (m, 3H); MS (EI) for $C_{23}H_{23}N_5OF_2$ HCl: 424 (MH^+).

[0641] 3-amino-6-(3-([(6-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 7.78 (m, 1H); 8.08 (s, 1H); 8.22 (m, 1H); 7.4 – 7.3 (m, 4H); 4.2 (m, 1H); 3.8 (m, 2H); 2.9 (m, 1H); 2.83 (d, 3H); 2.7 (m, 3H); 2.37 (m, 1H); 2.4 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OFBr$: 470 (MH^+).

[0642] 3-amino-6-(3-([(4-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 9.8 (br s, NH); 9.6 (br s, NH); 8.98 (m, 1H); 8.88 (s, 1H); 8.6 (s, 1H); 8.2 (m, 1H); 7.7 – 7.2 (m, 6H); 5.1 – 4.38 (m, 1H); 4.38 (m, 2H); 3.4 (m, 1H); 2.95 (m, 1H); 2.85 (d, 3H); 2.43 – 2.6 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OF$ HCl: 392 (MH^+).

[0643] 3-amino-6-(3-([(4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 9.1 (m, 1H); 8.8 (s, 1H); 8.6 (m, 1H); 8.4 (d, 1H); 8.74 (d, 1H); 7.5 – 7.2 (m, 3H); 4.6 (m, 1H); 3.9 (m, 2H); 3.4 (m, 1H); 3.0 (d, 3H); 2.9 (m, 1H); 2.5 (m, 1H); 1.3 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OF_2$: 410 (MH^+).

[0644] 3-amino-6-(3-([(6-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 8.6 (s, 1H); 8.0 (br s, 1H); 7.8 (s, 1H); 7.75 (m, 1H); 7.74 (m, 1H); 7.18 – 6.9 (m, 3H); 4.3 (m, 1H); 4.0 (m, 2H); 3.04 (d, 3H); 3.0 (m, 1H); 2.56 (m, 1H); 1.85 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OF$: 392 (MH^+).

[0645] 3-amino-6-(3-([(5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 8.6 (s, 1H); 8.0 (br s, 1H); 7.84 (s, 1H); 7.8 (d, 1H); 7.4 (m, 2H); 7.5 (m, 1H); 7.0 (m, 1H); 4.25 (t, 1H); 3.92

(m, 2H); 3.08 (d, 3H); 3.0 (m, 1H); 2.8 (m, 1H); 2.5 (m, 1H); 1.9 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OF_2$: 410 (MH^+).

[0646] 3-amino-6-(3-{[(5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $CDCl_3$): 8.64 (s, 1H); 8.4 (m, 1H); 7.88 (s, 1H); 7.76 (d, 1H); 7.4 (m, 2H); 6.78 (d, 1H); 6.64 (t, 1H); 4.6 (m, 1H); 3.9 (m, 2H); 3.2 (m, 1H); 3.1 (d, 3H); 2.85 (m, 1H); 2.4 (m, 1H); 2.1 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OF_2$: 410 (MH^+).

[0647] 3-amino-6-(3-{[(7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $DMSO-d_6$): 8.81 (s, 1H); 8.78 (br s, 1H); 8.1 (m, 1H); 8.05 (m, 1H); 7.4 (m, 3H); 7.02 (t, 1H); 4.2 (m, 1H); 3.9 (m, 2H); 3.4 (m, 1H); 2.85 (d, 3H); 2.76 (m, 1H); 2.4 (m, 1H); 1.8 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OFBr$: 472 (MH^+).

[0648] 3-amino-6-(3-{[(6-chloro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $DMSO-d_6$): 8.84 (s, 1H); 8.80 (m, 1H); 8.17 (s, 1H); 8.08 (d, 1H); 7.4 (m, 4H); 7.25 (m, 2H); 4.25 (m, 1H); 3.87 (m, 2H); 2.9 (m, 1H); 2.85 (d, 3H); 2.7 (m, 1H); 2.4 (m, 1H); 1.8 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OCl$: 408 (MH^+).

[0649] 3-amino-6-(3-{[(4,6-dichloro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $DMSO-d_6$): 8.84 (s, 1H); 8.8 (m, 1H); 8.1 (s, 1H); 8.06 (d, 1H); 7.39 (m, 4H); 4.2 (m, 1H); 3.87 (m, 2H); 2.9 (m, 1H); 2.85 (d, 3H); 2.77 (m, 1H); 2.4 (m, 1H); 1.8 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OCl_2$: 442 (MH^+).

[0650] 3-amino-6-(3-{[(4-chloro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $DMSO-d_6$): 8.84 (s, 1H); 8.80 (m, 1H); 8.2 (s, 1H); 8.05 (d, 1H); 7.4 – 7.23 (m, 4H); 4.25 (m, 1H); 3.87 (m, 2H); 2.95 (m, 1H); 2.85 (d, 3H); 2.78 (m, 1H); 2.4 (m, 1H); 1.84 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OCl$: 408 (MH^+).

[0651] 3-amino-6-(3-{[(6-bromo-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz; $DMSO-d_6$): 9.9 (m, 2H); 9.05 (m, 1H); 8.9 (s, 1H); 8.7 (s, 1H); 8.2 (d, 1H); 8.15 (s, 1H); 7.5 (m, 3H); 7.3 (d, 1H); 4.85 (m, 1H); 4.25 (m, 2H); 3.18 (m, 1H); 2.86 (m, 1H); 2.81 (d, 3H); 2.65 (m, 1H); 2.4 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OBr HCl$: 452 (MH^+).

[0652] 3-amino-6-(3-[[[(7-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide: MS (EI) for $C_{22}H_{22}N_5OFHCl$: 392 (MH^+).

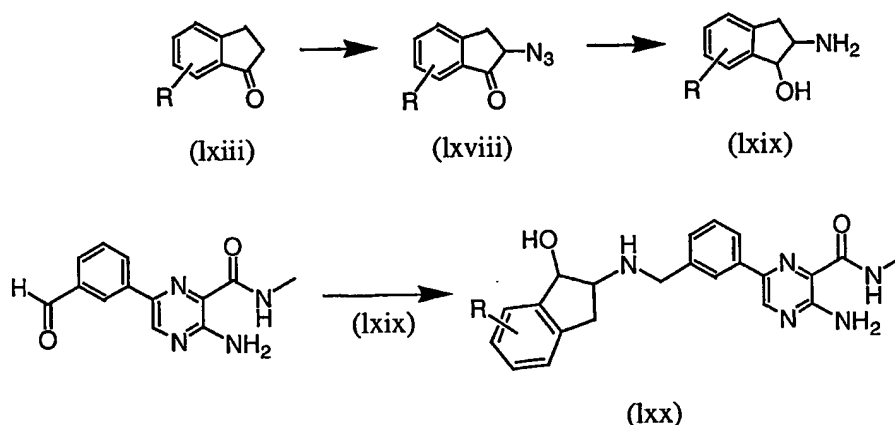
[0653] 3-amino-N-methyl-6-(3-[[[(2R)-1,2,3,4-tetrahydronaphthalen-2-ylamino]methyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.82 (s, 1H), 8.77 (dd, 1H), 8.08 (s, 1H), 8.02 (m, 1H), 7.62 (bs, 2H), 7.38 (m, 2H), 7.03 (s, 4H), 3.85 (s, 2H), 3.26 (m, 2H), 3.00 (dd, 1H), 2.86 (d, 3H), 2.81 (t, 1H), 2.68 (m, 1H), 2.56 (m, 1H), 2.02 (m, 1H), 1.56 (m, 1H); MS (EI) for $C_{23}H_{25}N_5O$: 388.4 (MH^+).

[0654] 3-amino-N-methyl-6-(3-[[[(2S)-1,2,3,4-tetrahydronaphthalen-2-ylamino]methyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.82 (s, 1H), 8.77 (dd, 1H), 8.11 (s, 1H), 8.04 (m, 1H), 7.62 (bs, 2H), 7.39 (m, 2H), 7.02 (s, 4H), 3.86 (s, 2H), 3.25 (m, 2H), 3.00 (dd, 1H), 2.88 (d, 3H), 2.81 (t, 1H), 2.72 (m, 1H), 2.62 (m, 1H), 2.04 (m, 1H), 1.58 (m, 1H); MS (EI) for $C_{23}H_{25}N_5O$: 388.4 (MH^+).

Example 32

[0655] Scheme 14 shows, analogously to Scheme 13 above, that 2-aminoindan-1-ols were also prepared for incorporation into compounds of the invention. Ketones (Ixxiii) were converted to azides (Ixxviii), followed by reduction to 2-aminoindan-1-ols (Ixxix). These amino alcohols were used to make compounds of the invention, for example compounds (Ixx), as depicted below.

Scheme 14



3-amino-6-(3-[[[(4-fluoro-1-hydroxy-2,3-dihydro-1H-inden-2-yl)amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide

- [0656] A solution of 4-fluoro-2,3-dihydro-1*H*-inden-1-one (1.29 g, 9.34 mmol) and HNIB (5.02 g, 12.1 mmol) in CH₃CN (190.0 mL) was refluxed for 1.5 h, until the solution was clear yellow. Upon cooling to r.t., NaN₃ (1.21 g, 18.7 mmol) was added and the solution was stirred at r.t. for 12 h. Additional NaN₃ (0.50 g, 7.7 mmol) was added and the solution stirred further at r.t. for 4 h. The reaction mixture was concentrated *in vacuo*, and the residue was taken up in CH₂Cl₂ (100 mL). This solution was washed with H₂O (75 mL), dried (Na₂SO₄), filtered and concentrated *in vacuo*. The residue was purified via column chromatography (SiO₂, 5:1 hexanes/EtOAc) to give 1.61 g (96% yield) of 2-azido-4-fluoro-2,3-dihydro-1*H*-inden-1-one: ¹H NMR (400 MHz, CDCl₃): 7.59 (d, 1H), 7.41 (m, 1H), 7.32 (t, 1H), 4.15 (dd, 1H), 3.55 (dd, 1H), 2.89 (dd, 1H).
- [0657] To a solution of 2-azido-4-fluoro-2,3-dihydro-1*H*-inden-1-one (0.81 g, 4.5 mmol) in MeOH (15.0 mL), was added NaBH₄ (0.19 g, 4.5 mmol) in one portion. After stirring for 10 min at r.t., the solution was concentrated *in vacuo*. The residue was taken up in EtOAc (100 mL), and this solution was washed with saturated NaHCO₃ (aq) (50 mL), dried (Na₂SO₄), filtered and concentrated *in vacuo* to give 0.64 g (73% yield) of 2-azido-4-fluoro-2,3-dihydro-1*H*-inden-1-ol as a white solid, which was used without further purification: ¹H NMR (400 MHz, CDCl₃): 7.23 (m, 2H), 6.97 (t, 1H), 5.15 (dd, 1H), 4.37 (q, 1H), 3.17 (d, 1H), 2.40 (d, 1H).
- [0658] A solution of 10% Pd/Carbon (0.070 g, 0.50 mmol)), in EtOAc (7.0 mL) was stirred under a H₂ (g) balloon for 2 h, after which time a solution of 2-azido-4-fluoro-2,3-dihydro-1*H*-inden-1-ol (0.98 g, 5.1 mmol) and Boc₂O (1.3 g, 6.1 mmol) in EtOAc (3.0 mL) was added. The mixture was stirred under a H₂ (g) balloon at r.t. for 12 h. Filtration through celite/fritted glass funnel and concentration *in vacuo* gave a crude residue, which was purified via column chromatography (SiO₂, 3:1 hexanes/EtOAc) to give 0.62 g (46% yield) of 1,1-dimethylethyl (4-fluoro-1-hydroxy-2,3-dihydro-1*H*-inden-2-yl)carbamate: ¹H NMR (400 MHz, CDCl₃): 7.18 (m, 2H), 6.94 (m, 1H), 5.21 (bs, 0.5H), 5.03 (bs, 0.5H), 4.37 (bs, 0.5H), 4.10 (q, 0.5H), 3.24-3.40 (m, 1H), 2.59-2.86 (m, 1H), 1.44 (s, 9H).
- [0659] To a solution of 1,1-dimethylethyl (4-fluoro-1-hydroxy-2,3-dihydro-1*H*-inden-2-yl)carbamate (0.30 g, 1.1 mmol) in CH₂Cl₂ (10 mL) was added TFA (1 mL), and the solution was heated to reflux. The solution was immediately cooled and concentrated *in vacuo*. The residue was taken up in MeOH and treated with Bio-Rad AG 1-X8 resin (hydroxide form) until pH 8. The product was filtered and concentrated *in vacuo* to give

0.12 g (64% yield) of 2-amino-4-fluoro-2,3-dihydro-1*H*-inden-1-ol, which was used without further purification for the subsequent reductive amination.

[0660] A solution of 3-amino-6-(3-formylphenyl)-*N*-methylpyrazine-2-carboxamide (0.15 g, 0.62 mmol), 2-amino-4-fluoro-2,3-dihydro-1*H*-inden-1-ol (0.12 g, 0.74 mmol), glacial AcOH (0.070 mL, 1.2 mmol) and NaBH(OAc)₃ (0.39 g, 1.8 mmol) in THF (2.5 mL) was heated to 70°C for 1 h. Upon cooling, saturated NaHCO₃ (aq) was added (25 mL). The aqueous layer was extracted with EtOAc (2 x 25 mL). The combined organic layers were dried (Na₂SO₄), filtered and concentrated *in vacuo*. The crude mixture was purified via HPLC (reverse-phase, CH₃CN/H₂O with 0.1% TFA). Upon removal of CH₃CN/H₂O, the product was taken up in MeOH and treated with Bio-Rad AG 1-X8 resin (hydroxide form) until pH 8. The product was filtered and concentrated *in vacuo*, to provide 14.1 mg (6% yield) of 3-amino-6-(3-([(4-fluoro-1-hydroxy-2,3-dihydro-1*H*-inden-2-yl)amino]methyl)phenyl)-*N*-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.81 (d, 1H), 8.77 (bs, 1H), 8.08 (d, 1H), 8.02 (d, 1H), 7.38 (m, 2H), 7.22 (m, 1H), 7.16 (d, 1H), 7.10 (d, 1H), 7.00 (q, 1H), 5.60 (d, 0.5H), 5.34 (bs, 0.5H), 4.91 (d, 0.5H), 4.84 (t, 0.5H), 3.84-4.02 (m, 2H), 3.18 (m, 1H), 2.98 (dd, 0.5H), 2.85 (d, 3H), 2.76 (dd, 0.5H), 2.52 (m, 2H); MS (EI) for C₂₂H₂₂N₅O₂F: 408 (MH⁺).

[0661] Using the analogous synthetic techniques beginning from (4-fluoro-2,3-dihydro-1*H*-inden-2-yl)amine [Haadsma-svensson, S.R. et al. WO 9 504 713, 1995], the following compound of the invention was prepared:

[0662] 3-amino-6-(3-([(4-fluoro-2,3-dihydro-1*H*-inden-2-yl)amino]methyl)phenyl)-*N*-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.81 (s, 1H), 8.75 (m, 1H), 8.06 (s, 1H), 8.02 (m, 1H), 7.37 (m, 2H), 7.14 (m, 1H), 7.02 (d, 1H), 6.91 (t, 1H), 3.83 (s, 2H), 3.60 (m, 1H), 3.11 (dd, 2H), 2.85 (d, 2H), 2.73-2.81 (m, 3H); MS (EI) for C₂₂H₂₂N₅OF: 392 (MH⁺).

Example 33

[0663] 3-amino-*N*-methyl-6-{3-[(naphthalen-2-ylamino)methyl]phenyl}pyrazine-2-carboxamide. A solution of 3-amino-6-(3-formylphenyl)-*N*-methylpyrazine-2-carboxamide (0.075 g, 0.29 mmol), naphthalen-2-amine (0.21 g, 1.5 mmol), glacial AcOH (0.033 mL, 0.56 mmol) and NaBH(OAc)₃ (0.093 g, 0.44 mmol) in THF (1.1 mL) was heated to 80°C for 12 h. Upon cooling, saturated NaHCO₃ (aq) was added (25 mL).

The aqueous layer was extracted with EtOAc (2 x 25 mL). The combined organic layers were dried (Na₂SO₄), filtered and concentrated *in vacuo*. The crude mixture was purified via HPLC (reverse-phase, CH₃CN/H₂O with 0.1% TFA). Upon removal of CH₃CN/H₂O, the product was taken up in MeOH and treated with Bio-Rad AG 1-X8 resin (hydroxide form) until pH 8. The product was filtered and concentrated *in vacuo*, to provide 29.1 mg (26% yield) of 3-amino-*N*-methyl-6-{3-[(naphthalen-2-ylamino)methyl]phenyl}pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.84 (s, 1H), 8.77 (d, 1H), 8.22 (s, 1H), 8.05 (dt, 1H), 7.62 (t, 2H), 7.51 (d, 1H), 7.41 (d, 2H), 7.27 (dt, 1H), 7.10 (dt, 2H), 6.77 (d, 1H), 6.57 (t, 1H), 4.46 (d, 2H), 2.85 (d, 3H); MS (EI) for C₂₃H₂₁N₅O: 384 (MH⁺).

[0664] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0665] 3-amino-*N*-methyl-6-{3-[(naphthalen-1-ylamino)methyl]phenyl}pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 8.78 (s, 1H), 8.70 (m, 1H), 8.25 (m, 1H), 8.20 (s, 1H), 7.99 (m, 2H), 7.73 (m, 1H), 7.42 (m, 2H), 7.36 (m, 2H), 7.16 (t, 1H), 7.05 (d, 1H), 6.96 (t, 1H), 6.42 (d, 1H), 4.57 (d, 2H), 2.84 (d, 3H); MS (EI) for C₂₃H₂₁N₅O: 384 (MH⁺).

Example 34

3-amino-6-[3-({[(3-fluorobiphenyl-4-yl)methyl]amino)methyl}phenyl]-*N*-methylpyrazine-2-carboxamide

[0666] A solution of 4-bromo-2-fluorobenzaldehyde (0.15 g, 0.74 mmol), tributyl(phenyl)stannane (0.27 mL, 0.81 mmol) and Pd(PPh₃)₄ (0.043 g, 0.037 mmol) in PhCH₃ (2.5 mL) was heated in a sealed reaction tube to 110°C for 12 h. Additional tributyl(phenyl)stannane (0.20 mL, 0.61 mmol) and Pd(PPh₃)₄ (0.043 g, 0.037 mmol) were added and the solution was further heated to 110°C for 2 h. The solvent was removed *in vacuo*. The residue was purified via column chromatography (SiO₂, 5:1 hexanes/EtOAc) to provide 0.23 g (>100% yield) of 3-fluorobiphenyl-4-carbaldehyde, which was used without further purification: MS (EI) for C₁₃H₉OF: 201 (MH⁺).

[0667] A solution of 3-fluorobiphenyl-4-carbaldehyde (0.030 g, 0.15 mmol), 3-amino-6-[3-(aminomethyl)phenyl]-*N*-methylpyrazine-2-carboxamide (0.038 g, 0.15 mmol), glacial AcOH (0.010 mL, 0.15 mmol) and NaBH(OAc)₃ (0.047 g, 0.22 mmol) in THF (0.6 mL) was stirred at r.t. for 12 h. Saturated NaHCO₃ (aq) was added (25 mL) and the aqueous

layer was extracted with EtOAc (2 x 25 mL). The combined organic layers were dried (Na_2SO_4), filtered and concentrated *in vacuo*. The crude mixture was purified via HPLC (reverse-phase, $\text{CH}_3\text{CN}/\text{H}_2\text{O}$ with 0.1% TFA). Upon removal of $\text{CH}_3\text{CN}/\text{H}_2\text{O}$, the product was taken up in MeOH and treated with Bio-Rad AG 1-X8 resin (hydroxide form) until pH 8. The product was filtered and concentrated *in vacuo*, to provide 13.4 mg (20% yield) of 3-amino-6-[3-({[(3-fluorobiphenyl-4-yl)methyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.84 (s, 1H), 8.79 (d, 1H), 8.09 (bs, 1H), 8.05 (m, 1H), 7.70 (m, 2H), 7.61 (t, 1H), 7.36-7.53 (m, 8H), 3.80 (d, 4H), 2.86 (d, 3H); MS (EI) for $\text{C}_{26}\text{H}_{24}\text{N}_5\text{OF}$: 442 (MH^+).

[0668] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0669] 3-amino-6-[3-({[(2-fluoro-4-(2-thienyl)phenyl)methyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.83 (s, 1H), 8.79 (m, 1H), 8.07 (bs, 1H), 8.05 (m, 1H), 7.58 (m, 2H), 7.54 (d, 1H), 7.37-7.50 (m, 4H), 7.15 (m, 1H), 3.78 (d, 4H), 3.17 (d, 1H), 2.86 (d, 3H); MS (EI) for $\text{C}_{24}\text{H}_{22}\text{N}_5\text{OFS}$: 448 (MH^+).

[0670] 3-amino-6-[3-({[(2-fluoro-4-furan-2-ylphenyl)methyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.81 (s, 1H), 8.77 (m, 1H), 8.06 (bs, 1H), 8.02 (m, 1H), 7.74 (d, 1H), 7.35-7.57 (m, 4H), 7.00 (d, 1H), 6.59 (m, 1H), 3.75 (d, 4H), 2.84 (d, 3H); MS (EI) for $\text{C}_{24}\text{H}_{22}\text{N}_5\text{O}_2\text{F}$: 432 (MH^+).

[0671] 3-amino-6-[3-({[(2-fluoro-5-(2-thienyl)phenyl)methyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.80 (s, 1H), 8.75 (m, 1H), 8.07 (bs, 1H), 8.02 (m, 1H), 7.77 (dd, 1H), 7.55 (m, 1H), 7.50 (dd, 1H), 7.37-7.44 (m, 3H), 7.18 (dd, 1H), 7.10 (dd, 1H), 3.80 (d, 4H), 3.16 (d, 1H), 2.83 (d, 3H); MS (EI) for $\text{C}_{24}\text{H}_{22}\text{N}_5\text{OFS}$: 448 (MH^+).

[0672] 3-amino-6-[3-({[(4-ethenyl-2-fluorophenyl)methyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 8.80 (s, 1H), 8.75 (m, 1H), 8.04 (bs, 1H), 8.01 (m, 1H), 7.25-7.48 (m, 5H), 6.69 (dd, 1H), 5.86 (d, 1H), 5.27 (d, 1H), 3.75 (d, 4H), 3.16 (d, 1H), 2.85 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{22}\text{N}_5\text{OF}$: 392 (MH^+).

Example 35

2-amino-5-[3-hydroxymethyl]phenyl]-N-methylpyridine-3-carboxamide

[0673] 2-Amino-5-bromopyridine-3-carboxylic acid was prepared by bromination, para- to the 2-amino group of commercially available 2-aminonicotinic acid as described in US patents 3,950,160 and 4,361,700.

[0674] Methyl 2-amino-5-bromopyridine-3-carboxylate: A slurry of 2-amino-5-bromopyridine-3-carboxylic acid (7 g, 32.2 mmol) in methanol (150 mL) and H₂SO₄ (7 mL) was heated to reflux for 48 h. The reaction mixture was concentrated to ca. 40 mL. Water (100 mL) was added to the residual oil, and the mixture neutralized with sodium carbonate. Ethyl acetate (400 mL) was added and the organic layer was washed with saturated aqueous sodium chloride then dried (Na₂SO₄) and concentrated to give the title compound (5.5 g, 73.9% yield).

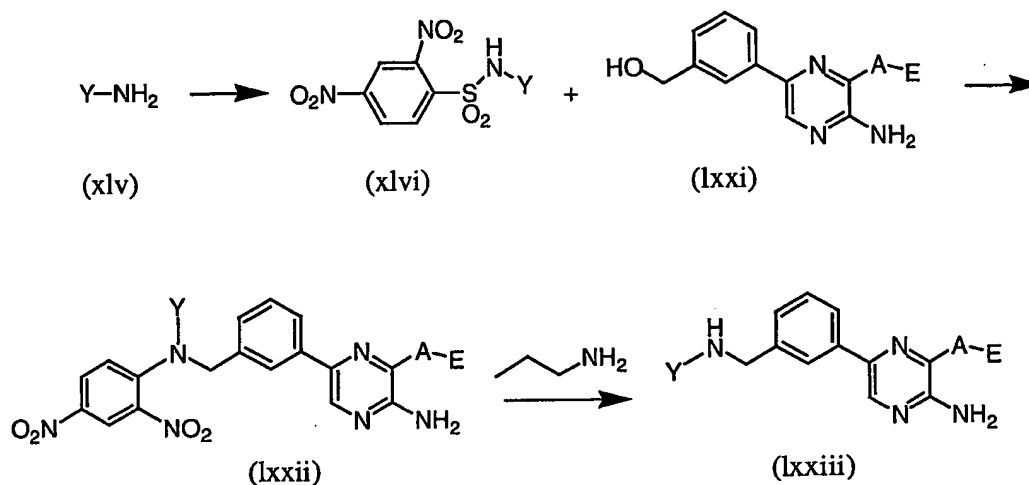
[0675] 2-Amino-5-bromopyridine-N-methylpyridine-3-carboxamide: A slurry of the above methyl 2-amino-5-bromopyridine-3-carboxylate (3.0 g, 13 mmol) in methanol was cooled to 0 °C, and methylamine gas was passed through the solution at a moderate rate for 5 min. The mixture was stirred in sealed tube at 50 °C for 24 h, and the methanol was removed in vacuo to provide 2-amino-5-bromopyridine-N-methylpyridine-3-carboxamide as a white solid (2.5 g, 80.6% yield): ¹H NMR (400 MHz; DMSO-d₆): 8.54 (br s, 1H); 8.19 (d, 1H); 8.08 (d, 1H); 7.25 (s, 1H); 2.72 (d, 3H).

[0676] 2-Amino-5-[3-hydroxymethyl]phenyl]-N-methylpyridine-3-carboxamide: To a solution of 2-amino-5-bromopyridine-N-methylpyridine-3-carboxamide (0.15 g, 0.66 mmol) in DMF (10 mL) was added 3-hydroxymethylphenyl boronic acid (129 mg, 0.85 mmol), Pd(dppf)₂Cl₂ (10 mol% yield) and sodium carbonate (0.3 g) and this mixture was stirred under a nitrogen atmosphere for 5 minutes. The mixture was heated at 85 °C for 4 h. The reaction mixture was cooled to room temperature. The mixture was extracted with ethyl acetate (3x10 mL) and the combined organic fractions were sequentially washed with aqueous ammonium chloride (2x5 mL) and brine (2x5 mL), dried over anhydrous sodium sulfate, filtered and evaporated in vacuo. The residue was triturated with diethyl ether/hexanes (2:1) and the solid was collected by vacuum filtration. Purification by column chromatography afforded a white solid (100 mg, 59% yield): ¹H NMR (400 MHz; DMSO-d₆): 8.6 (br s, NH); 8.4 (d, 1H); 8.2 (d, 1H); 7.6 (s, 1H); 7.55 (d, 1H); 7.4 (t, 1H); 7.27 (d, 1H); 7.2 (br s, 2NH); 5.25 (t, 1H); 4.55 (d, 2H); 2.80 (d, 3H); MS (EI) for C₁₄H₁₅N₃O: 258 (MH⁺).

Example 36

[0677] Scheme 15 depicts a general synthetic route for compounds of the invention having substituents -W-X-Y according to formula I, wherein, for example, W is phenylene, X is -CH₂NH-, and Y is various groups. Note, as defined above any of W, X, or Y are optionally substituted; this is only a set of examples. Analogous to Scheme 10, primary amine (xlv), which in this example incorporates Y, is incorporated into a compound of the invention, (lxxiii). For example, amine (xlv), which has Y as part of its structure for example, is converted to 2,4-dinitrosulfonamide (xlvi). Sulfonamide (xlvi) is used to make sulfonamide (lxxii). The transformation (xlvi) to (lxxii) is carried out under Mitsunobu conditions, in this case where the alcohol partner is intermediate, compound of the invention, (lxxi). As described above in relation to Scheme 10, the sulfonamide bond is cleaved, typically but not necessarily via addition of a primary amine e.g. *n*-propyl amine, to free secondary amine (lxxiii), which in this case is a compound of the invention.

Scheme 15



2-Amino-5-(3-[[[(2,4-dinitrophenyl)sulfonyl][(1S)-2,3-dihydro-1H-inden-1-yl]amino]methyl]phenyl]-N-methylpyridine-3-carboxamide

[0678] N-[(1S)-2,3-dihydro-1H-inden-1-yl]-2,4-dinitrobenzenesulfonamide: To a stirred, ice-cooled solution of 4.00 g of (S)-(+)-1-aminoindan (30.0 mmol, from Sigma-Aldrich) in 113 mL of THF was added 9.60 g (36.0 mmol, 1.20 eq.) of 2,4-dinitrobenzenesulfonyl chloride, followed by 7.80 mL (44.8 mmol, 1.49 eq.) of N,N-diisopropylethylamine. The ice bath was removed, and the mixture was stirred at room temperature for 19 h and then concentrated. The residue was taken up in EtOAc, washed with 2 x H₂O, dried over Na₂SO₄, filtered, and concentrated. The residue was sonicated in ca. 30 mL of EtOAc

until a precipitate was formed. The mixture was diluted to ca. 150 mL with hexanes. Filtration afforded pure product as a tan solid (8.59 g, 78.8% yield).

[0679] Triphenylphosphine (6.9 g, 26 mmol) was added to a stirred solution of 2-amino-5-[3-(hydroxymethyl)phenyl]-N-methylpyridine-3-carboxamide (5.16 g, 20 mmol) and N-[(1S)-2,3-dihydro-1H-inden-1-yl]-2,4-dinitrobenzenesulfonamide (9.4 g, 26 mmol) in benzene (150 mL). Diethyl azodicarboxylate (4.5 g, 26 mmol) was then added drop wise and mixture was stirred at ambient temperature for 4 h. Ethyl acetate (400 mL) was added and the organic layer was washed with saturated aqueous sodium chloride then dried (Na_2SO_4). Filtration and concentration followed by column chromatography (ethyl acetate/hexane, 9:1) gave desired product (8.2 g, 50 % yield).

[0680] 2-amino-5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-methylpyridine-3-carboxamide: 2-Amino-5-(3-{[(2,4-dinitrophenyl)sulfonyl][(1S)-2,3-dihydro-1H-inden-1-yl]amino}methyl}phenyl)-N-methylpyridine-3-carboxamide was treated with excess n-propylamine (4 eq) in CH_2Cl_2 (150 mL). The mixture was diluted with additional ethyl acetate (200 mL). The mixture was subsequently washed with aqueous saturated sodium carbonate (3 x 30 mL) and brine (3 x 30 mL). The combined organic fractions were dried over anhydrous sodium sulfate, filtered and concentrated in vacuo. Purification of the residue by preparative HPLC (acetonitrile/water) gave desired product (5.8 g). The compound was converted to the hydrochloride salt using 4 M solution HCl in dioxane: ^1H NMR (400 MHz; DMSO-d_6): 9.9 (br s, NH); 9.02 (s, 1H); 8.9 (s, 1H); 8.6 (s, 1H); 8.35 (br s, NH); 8.2 (s, 1H); 7.8 (d, 2H); 7.75 (m, 2H); 7.37 (m, 2H); 7.29 (m, 1H); 4.85 (m, 1H); 4.25 (m, 2H); 3.4 (m, 1H); 2.9 (m, 1H); 2.80 (d, 3H); 2.43 (m, 1H); 2.4 (m, 1H); MS (EI) for $\text{C}_{23}\text{H}_{24}\text{N}_4\text{O HCl}$: 374 (MH^+).

[0681] 2-amino-5-(3-{[(6-bromo-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyridine-3-carboxamide: ^1H NMR (400 MHz; DMSO-d_6): 9.8 (br s, 2H); 9.15 (br s, 1H); 8.8 (s, 1H); 8.31 (s, 1H); 8.08 (s, 1H); 7.85 (d, 1H); 7.6 (m, 2H); 7.3 (d, 1H); 4.85 (m, 1H); 4.3 (m, 2H); 3.2 (m, 1H); 2.85 (m, 1H); 2.81 (d, 3H); 2.56 (m, 1H); 2.4 (m, 1H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_4\text{OBr HCl}$: 451 (MH^+).

[0682] 2-amino-5-(3-{[(4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyridine-3-carboxamide: ^1H NMR (400 MHz; DMSO-d_6): 10.0 (br s, 2H); 9.18 (m, 1H); 8.8 (s, 1H); 8.6 (s, 1H); 8.30 (br s, 2H); 8.25 (s, NH); 7.85 - 7.2 (m, 5H); 4.85 (m, 1H); 4.4 (m, 2H); 3.2 (m, 1H); 2.9 (m, 1H); 2.85 (d, 3H); 2.6 - 2.4 (m, 2H); MS (EI) for $\text{C}_{23}\text{H}_{22}\text{N}_4\text{OF}_2 \text{ HCl}$: 409 (MH^+).

[0683] 2-amino-5-(3-{[(7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyridine-3-carboxamide: MS (EI) for $C_{23}H_{22}N_4OFBr$ HCl: 469 (MH^+).

[0684] 2-amino-5-(3-{[(4-chloro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyridine-3-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 9.4 (br s, NH); 8.9 (br s, 1H); 8.65 (br s, 1H); 8.6 (s, 1H); 8.2 (s, 1H); 7.75 (m, 2H); 7.6–7.48 (m, 3H); 7.7 (d, 1H); 4.85 (m, 1H); 4.3 (m, 2H); 3.8 (m, 1H); 3.18 (m, 1H); 2.85 (m, 3H); 2.59–2.4 (m, 2H); MS (EI) for $C_{23}H_{23}N_4OCl$ HCl: 407 (MH^+).

[0685] 2-amino-5-(3-{[(5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyridine-3-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 9.83 (br s, NH); 9.09 (br s, NH); 8.82 (s, 1H); 8.58 (s, 1H); 8.22 (s, 1H); 7.99 (m, 1H); 7.8 (m, 1H); 7.67–7.44 (m, 2H); 4.85 (s, 1H); 4.27 (m, 2H); 3.2 (m, 1H); 2.95 (m, 1H); MS (EI) for $C_{23}H_{22}N_4OF_2$: 408 (MH^+).

[0686] 2-amino-5-(3-{[(5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyridine-3-carboxamide: 1H NMR (400 MHz; DMSO- d_6): 9.7 (br s, NH); 9.4 (br s, 1H); 9.0 (br s, 1H); 8.7 (s, 1H); 8.57 (s, 1H); 8.1 (s, 1H); 7.8 (m, 1H); 7.5 (m, 1H); 7.2 (m, 2H); 4.85 (m, 1H); 4.3 (m, 2H); 3.89 (m, 1H); 3.18 (m, 1H); 2.85 (m, 3H); 2.69–2.5 (m, 2H); MS (EI) for $C_{23}H_{22}N_4OF_2$ HCl: 409 (MH^+).

[0687] Using a strategy that combines synthetic technique as described in relation to both Scheme 14 and Scheme 15, the following compounds were made:

3-Amino-6-[3-({[6-(3-hydroxypropyl)-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide:

[0688] 5-Bromo-2,3-dihydro-1H-inden-1-one (1.68g, 7.99 mmol) was dissolved in DMF and treated with triethylamine (1.61 g, 15.98 mmol), copper iodide (152 mg, 0.79 mmol) and prop-2-yn-1-ol (2.24 g, 39.96 mmol) in the presence of $Pd(PPh_3)_4$ (462 mg, 0.39 mmol) at ambient temperature. After being refluxed for 16 h, the reaction mixture was extracted with ethyl acetate (100 mLx3) and washed with brine. The combined organic layers were dried over $MgSO_4$. Solvent removal under vacuum gave crude tan oil. Purification by preparative thin-layer chromatography afforded 5-(hydroxyprop-1-yn-1-yl)-2,3-dihydro-1H-indene-1-one (1.06 g, 72 % yield): 1H NMR (400 MHz, CD_3OD): δ 7.63 (d, 1H), 7.59 (s, 1H), 7.42 (d, 1H), 4.45 (s, 2H), 3.18 (m, 2H), 2.69 (m, 2H); MS (EI) for $C_{12}H_{10}O_2$: 187.20 (MH^+).

- [0689] To a solution of 5-(hydroxyprop-1-yn-1-yl)-2,3-dihydro-1*H*-indene-1-one (1.06g, 5.71 mmol) in MeOH was added 10% Pd/C and hydrogenated by means of Parr shaker. After 12 h, the reaction mixture was filtered on Celite and concentrated in vacuo to give 5-(3-hydroxypropyl)-2,3-dihydro-1*H*-inden-1-one (1.03 g, 94 % yield): MS (EI) for C₁₂H₁₄O₂: 191.24 (MH⁺).
- [0690] 5-(Hydroxyprop-1-yn-1-yl)-2,3-dihydro-1*H*-indene-1-one (1.03 g, 5.42 mmol) was dissolved in DMF and treated with chloro-tert-butyldimethylsilane (981 mg, 6.51 mmol), imidazole (443 mg, 6.51 mmol) at 0°C. The reaction mixture was allowed to reach ambient temperature. After 10 h, the reaction mixture was extracted with ethyl acetate (100 mL x 3), washed with water, brine and dried over MgSO₄. The organic layer was concentrated under vacuum and purified by flash column chromatography to yield the 5-(3-[[[(1,1-dimethylethyl)(dimethyl)silyl]oxy]propyl]-2,3-dihydro-1*H*-inden-1-one (1.54 g, 94 % yield): ¹H NMR (400 MHz, CDCl₃): δ 7.63 (d, 1H), 7.59 (s, 1H), 7.42 (d, 1H), 4.45 (t, 2H), 3.10 (m, 2H), 2.78 (m, 2H), 2.64 (m, 2H), 1.82 (m, 2H), 0.95 (s, 9H), 0.06 (s, 6H).
- [0691] 5-(3-[[[(1,1-Dimethylethyl)(dimethyl)silyl]oxy]propyl]-2,3-dihydro-1*H*-inden-1-one (1.54 g, 5.07 mmol) was dissolved in MeOH (20 mL) and treated with NaBH₄ (231 mg, 6.10 mmol) at 0°C. After 1 h, the reaction mixture was quenched by slow addition of water at 0°C. The reaction mixture was extracted with ethyl acetate (100 mL x 3), washed with water, brine and dried over MgSO₄. The organic layer was concentrated under vacuum and moved to the next reaction without further purification. To a solution of alcohol (1.56 g, 5.079 mmol) were added DPPA (1.81 g, 6.60 mmol), DBU (1.00 g, 6.60 mmol) at 0°C. The reaction mixture was allowed to reach ambient temperature and stirred for 10 h. The reaction mixture was concentrated in vacuo and purified by flash column chromatography to give 1-[5-(3-[[[(1,1-dimethylethyl)(dimethyl)silyl]oxy]propyl]-2,3-dihydro-1*H*-inden-1-yl]-2λ⁵-triaz-1-en-2-yne (1.05 g, 63 %, two steps): ¹H NMR (400 MHz, CDCl₃): δ 7.25 (d, 1H), 7.05 (m, 2H), 4.79 (m, 1H), 3.59 (t, 2H), 3.00 (m, 1H), 2.78 (m, 1H), 2.63 (m, 2H), 2.38 (m, 1H), 2.06 (m, 1H), 1.77 (m, 2H), 1.88 (s, 9H), 0.05 (s, 6H).
- [0692] 1-[5-(3-[[[(1,1-Dimethylethyl)(dimethyl)silyl]oxy]propyl]-2,3-dihydro-1*H*-inden-1-yl]-2λ⁵-triaz-1-en-2-yne was hydrogenated in MeOH on a Parr shaker (30 psi) overnight and filtered on Celite. The organic layer was concentrated and dissolved in THF (15 mL). To a solution of resulting amine were added 3,5-dinitrobenzenesulfonyl

chloride (1.02 g, 3.81 mmol), diisopropylethylamine (616 mg, 4.76 mmol) at 0°C. The reaction mixture was allowed to reach to ambient temperature and stirred for 3 h to complete the reaction. The reaction mixture was concentrated in vacuo and purified by flash column chromatography to give *N*-[5-(3-([(1,1-dimethylethyl)(dimethyl)silyl]oxy)propyl)-2,3-dihydro-1*H*-inden-1-yl]-3,5-dinitrobenzenesulfonamide (1.16 g, 68 % yield, two steps).

[0693] To a solution of 3-amino-6-[3-(hydroxymethyl)phenyl]-*N*-methylpyrazine-2-carboxamide, 403 mg, 1.55 mmol) were added *N*-[5-(3-([(1,1-dimethylethyl)(dimethyl)silyl]oxy)propyl)-2,3-dihydro-1*H*-inden-1-yl]-3,5-dinitrobenzenesulfonamide (950 mg, 1.77 mmol), triphenylphosphine (466 mg, 1.77 mmol), diethyl azodicarboxylate (309 mg, 1.77 mmol) at 0 °C and the reaction mixture was allowed to reach ambient temperature. After 10 h, the reaction mixture was extracted with ethyl acetate (100 mL x 3), washed with water, brine and dried over MgSO₄. The organic layer was concentrated in vacuo and purified by flash column chromatography to yield 3-amino-6-[3-([(5-(3-([(1,1-dimethylethyl)(dimethyl)silyl]oxy)propyl)-2,3-dihydro-1*H*-inden-1-yl][(3,5-dinitrophenyl)sulfonyl]amino)methyl)phenyl]-*N*-methylpyrazine-2-carboxamide (655 mg, 54% yield): ¹H NMR (400 MHz, CDCl₃): δ 8.45 (s, 1H), 8.28 (d, 1H), 8.02 (m, 1H), 7.91 (m, 1H), 7.92 (d, 1H), 7.58 (m, 2H), 7.21 (m, 2H), 7.06 (m, 3H), 5.74 (t, 1H), 4.45 (d, 1H), 4.27 (d, 1H), 3.61 (m, 2H), 3.08 (d, 3H), 2.88 (m, 2H), 2.65 (m, 3H), 2.12 (m, 1H), 1.78 (m, 2H), 0.89 (s, 9H), 0.05 (s, 6H).

[0694] 3-Amino-6-[3-([(5-(3-([(1,1-dimethylethyl)(dimethyl)silyl]oxy)propyl)-2,3-dihydro-1*H*-inden-1-yl][(3,5-dinitrophenyl)sulfonyl]amino)methyl)phenyl]-*N*-methylpyrazine-2-carboxamide (600 mg, 0.77 mmol) was dissolved in THF and refluxed for an hour in the presence of tetrabutylammonium fluoride in THF. The reaction mixture was partitioned with water and extracted with ethyl acetate (10 mL x 3) and dried over MgSO₄. Filtration and concentration under vacuum gave tan oil, which was subjected to the solution of dichloromethane and isopropylamine. After being stirred for 30 min, the solvent was removed in vacuo and purified by flash column chromatography to give the 3-amino-6-[3-([(6-(3-hydroxypropyl)-2,3-dihydro-1*H*-inden-1-yl]amino)methyl)phenyl]-*N*-methylpyrazine-2-carboxamide (110 mg, 33% yield): ¹H NMR (400 MHz, CD₃OD): δ 8.59 (s, 1H), 8.06 (bs, 1H), 7.91 (s, 1H), 7.52 (m, 1H), 7.40 (m, 2H), 7.11 (d, 1H), 7.08 (s, 1H), 7.03 (d, 1H), 4.32 (d, 1H), 3.66 (t, 2H), 3.02 (d, 3H), 2.99 (m, 1H), 2.81 (m, 1H),

2.69 (t, 2H), 2.44 (m, 1H), 1.90 (m, 4H), 1.32 (m, 3H), 0.82 (m, 2H); MS (EI) for $C_{25}H_{29}N_5O_2$: 432.23 (MH^+).

[0695] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0696] 3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)pyrazine-2-carbonitrile: 1H NMR (400 MHz, d_6 -DMSO): 8.93 (s, 1H), 7.98 (s, 1H), 7.81 (d, 1H), 7.46 (br s, 2H), 7.44 – 7.38 (m, 3H), 7.22 (m, 1H), 7.17 (m, 2H), 4.16 (t, 1H), 3.86 (m, 2H), 2.94 (m, 1H), 2.73 (m, 1H), 2.32 (m, 1H), 1.82 (m, 1H); MS (EI) for $C_{21}H_{19}N_5$: 342.2 (MH^+).

Example 37

5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine

[0697] N-{3-[5-amino-6-(5-benzyl-4H-1,2,4-triazole-3-yl) pyrazin-2-yl]benzyl}-2,4-dinitro-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzene sulfonamide: Triphenylphosphine (47 mg, 0.17 mmol) was added to a stirred solution of {3-[5-amino-6-(5-phenyl-4H-1,2,4-triazol-3-yl) pyrazine-2-yl] phenyl}methanol (50 mg, 0.14 mmol) and N-[(1S)-2,3-dihydro-1H-inden-1-yl]-2,4-dinitrobenzen sulfonamide (60 mg, 0.16 mmol) in THF. Diethyl azodicarboxylate (31 mg, 0.17 mmol) was then added and mixture was stirred at ambient temperature 4 h. Ethyl acetate (20 mL) was added, and the organic layer was washed with saturated aqueous sodium chloride, and then dried (Na_2SO_4). Filtration and concentration followed by chromatography on silica gel afforded the title compound: MS (EI) for $C_{35}H_{29}N_9O_6S$: 704 (MH^+).

[0698] The solution of crude benzene sulfonamide (50 mg) in dichloromethane (3 mL) was treated with excess n-propylamine (4 eq). The solvent was evaporated, and the residue chromatographed over silica gel with EtOAc-hexane (7:1) affording 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine as a yellow solid (15 mg, 22% yield); 1H NMR (400 MHz; $DMSO-d_6$): 8.78 (s, 1H); 8.18 (s, 1H); 8.16 (d, 1H); 7.65 (br s, NH); 7.48 – 7.1 (m, 11H); 4.2 (m, 3H); 3.8 (m, 2H); 2.9 (m, 1H);); 2.7 (m, 1H); 2.3 (m, 1H);); 1.8 (m, 1H); MS (EI) for $C_{29}H_{27}N_7$: 474 (MH^+).

[0699] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-(4H-1,2,4-triazol-3-yl)pyrazin-2-amine: ^1H NMR (400 MHz; DMSO- d_6): 9.9 (m, 1H); 8.88 (s, 1H); 8.68 (s, 1H); 8.4 (br s, 1H); 8.24 (d, 1H); 7.85 (d, 1H); 7.6 (m, 2H); 7.4 – 7.2 (m, 3H); 4.8 (m, 1H); 4.4 (m, 2H); 3.2 (m, 1H); 2.9 (m, 1H); 2.55 (m, 1H); 2.4 (m, 1H); MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_7$ HCl: 384 (MH^+).

[0700] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine: ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.6 (s, 1H); 8.4 (d, 1H); 7.8 (d, 1H); 7.6 – 7.25 (m, 5H); 4.48 (s, 1H); 4.4 (m, 2H); 3.2 (m, 1H); 2.9 (m, 1H); 2.5 (m, 1H); 2.35 (m, 1H); MS (EI) for $\text{C}_{26}\text{H}_{29}\text{N}_7$ HCl: 440 (MH^+).

[0701] (1S)-N-{[3-(6-{5-[(methyloxy)methyl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)phenyl]methyl}-2,3-dihydro-1H-inden-1-amine: ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.15 (s, 1H); 8.07 (d, 1H); 7.7 (br s, 1H); 7.47 (t, 1H); 7.4 (t, 1H); 5.25 (t, 1H); 4.6 (d, 2H); 3.1 – 2.9 (m, 3H); 2.4 (m, 2H); 2.2 – 1.9 (m, 3H); 1.8 – 1.6 (m, 3H); 1.04 (t, 3H); MS (EI) for $\text{C}_{20}\text{H}_{25}\text{N}_7\text{O}$: 380 (MH^+).

[0702] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine: ^1H NMR (400 MHz; DMSO- d_6): 10.4 (br s, NH); 9.98 (br s, NH); 8.9 (s, 1H); 8.77 (s, 1H); 8.3 (d, 1H); 7.84 (d, 1H); 7.8 (br s, 2NH); 7.5 (m, 2H); 7.3 (m, 3H); 4.8 (m, 1H); 4.3 (m, 2H); 3.8 – 3.6 (m, 3H); 3.4 – 2.9 (m, 6H); 2.55 – 1.6 (m, 6H); 1.3 (t, 3H); MS (EI) for $\text{C}_{29}\text{H}_{34}\text{N}_8$ HCl: 495 (MH^+).

Example 38

3-Amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)pyrazine-2-carboxamide

[0703] A solution methyl 3-amino-6-[3-(hydroxymethyl)phenyl] pyrazine-2-carboxylate (37.4 mg, 0.1 mmol) in methanol (5 mL) was cooled to 0 $^{\circ}\text{C}$, and ammonia gas was passed through the solution at a moderate rate for 2 min. The mixture was stirred in sealed tube at 50 $^{\circ}\text{C}$ for 24 h then the methanol was removed in vacuo to provide 2-amino-5-N-methylpyrazine-3-carboxamide as a solid (20 mg, 55.7% yield): ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.3 (br s, 1H); 8.15 (s, 1H); 8.0 (d, 1H); 7.7 (s, 1H); 7.6 (br s, 2H); 7.4 (m, 3H); 7.2 (m, 3H); 4.2 (m, 1H); 3.89 (m, 2H); 2.9 (m, 1H); 2.7 (m, 1H); 2.3 (m, 1H); 1.8 (m, 1H); MS (EI) for $\text{C}_{21}\text{H}_{21}\text{N}_5\text{O}$: 360 (MH^+).

[0704] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl})phenyl)-N-(2,2,2-trifluoroethyl)pyrazine-2-carboxamide was prepared from methyl 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carboxylate and 2,2,2-trifluoroethylamine in a manner analogous to that described in the previous example. ¹H NMR (400 MHz; DMSO-d₆): 9.9 (br s, 1H); 9.88 (br s, 1H); 9.5 (t, 1H); 8.97 (s, 1H); 8.75 (s, 1H); 8.24 (m, 1H); 7.84 (d, 1H); 7.5 (m, 2H); 7.4 – 7.3 (m, 3H); 4.8 (m, 1H); 4.3 (m, 2H); 4.2 (m, 2H); 3.4 (m, 1H); 2.9 (m, 1H); 2.5 (m, 1H); 2.4 (m, 1H); MS (EI) for C₂₃H₂₂N₅OF₃ HCl: 442 (MH⁺).

Example 39

3-Amino-6-bromopyrazine-2-carboxamide

[0705] A suspension of methyl 3-aminopyrazine-2-carboxylate (5 g, 32.6 mmol) and 150 mL of dry methanol saturated with anhydrous ammonia were heated in an autoclave at 110-115 °C for 24 h. Concentration and filtration afforded 3-aminopyrazine-2-carboxamide (4.4 g, 97.7% yield) as a yellow solid.

[0706] To a solution 3-aminopyrazine-2-carboxamide (3 g, 22 mmol) in acetic acid (25 mL) was drop wise added bromine (4.2 g, 26.5 mmol) at room temperature the reaction mixture was stirred for 0.5 h and Na₂CO₃ (2 eq) was added slowly. Upon added of the reaction, the solvent was removed in vacuo, water (50 mL) was added to the residue and the precipitate was collected on a filter and washed with hexane (50 mL) to give 3-amino-6-bromopyrazine-2-carboxamide (4.5g) as a yellow solid: MS (EI) for C₅H₅BrN₄O: 217 (MH⁺).

[0707] 3-Amino-6-bromopyrazine-2-carbonitrile: 3-Amino-6-bromopyrazine-2-carboxamide (1.5 g, 7 mmol) was dissolved in pyridine (15 mL). The solution was then cooled in an ice bath and POCl₃ (2 mL) was drop wise added. The resulting mixture was stirred at room temperature for 2.5 h. The solvent was removed and the residue was quenched by the careful addition of ice cooled Na₂CO₃ (5 mL). Ethyl acetate was added (45 mL) and the organic layer was separated, washed with brine (5 mL), HCl (0.1 M aqueous solution, 1 mL) and water (5 mL), dried with K₂CO₃ and concentrated. The crude material was triturated with hexane, and the resulting tan solid was filtered to provide 0.8g (58% yield) of the title compound: ¹H NMR (400 MHz; DMSO-d₆): 8.42 (s, 1H); 7.60 (br s, NH); MS (EI) for C₅H₃BrN₄: 199 (MH⁺).

[0708] 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carbonitrile: A mixture of 3-amino-6-bromopyrazine-2-carbonitrile (7.7 g, 37.9 mmol), 3-hydroxymethylphenyl boronic acid (7.4 g, 49.3 mmol), Pd(dppf)₂Cl₂ (10 mol% yield), and triethylamine (16.6 mL, 113 mmol) in DMF (50 mL) were heated at 85 °C for 12 h. The reaction was cooled to room temperature and water (50 mL), EtOAc (100 mL) was added. The organic layer was washed with brine (50 mL), dried (Na₂SO₄) and passed through a plug of Celite. The solvent was removed and the residue triturated with ether (40 mL) to give a title compound (6 g) as a white solid: ¹H NMR (400 MHz; DMSO-d₆): 8.9 (s, 1H); 7.9 (s, 1H); 7.8 (d, 1H); 7.5 – 7.3 (m, 4H); 5.27 (t, 1H); 4.15 (d, 1H); MS (EI) for C₁₂H₁₀N₄O: 227 (MH⁺).

Example 40

Preparation of Triazoles

[0709] {3-[5-amino-6-(5-phenyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol: A mixture of 3-amino-6-[3-(hydroxymethyl) phenyl]pyrazine-2-carbonitrile (0.11 g, 0.5 mmol) and benzoic hydrazide (0.15 g, 1 mmol) were heated at 180 – 200 °C for 30 min. After cooling, the reaction mass was taken up with 5mL DMF and the obtained solution was diluted with EtOAc and water, clarified with charcoal and filtered on Celite. The organic layer was separated. After drying over sodium sulfate, the ethyl acetate was evaporated under reduced pressure. Purification of the residue by preparative HPLC (Acetonitrile/water) gave desired product (50 mg): ¹H NMR (400 MHz; DMSO-d₆): 8.3 (s, 1H); 8.1 (m, 2H); 7.6 - 7.3 (m, 7H); 5.45 (t, 1H); 4.6 (d, 2H); MS (EI) for C₂₃H₂₂N₅OF₃ HCl: 442 (MH⁺).

[0710] 1-ethyl piperidine-3-carbohydrazide: Using the procedure (as described in J. Med. Chem. (1990) 33 (1), 311-317), N-alkylation of ethyl nipecotate with 1-bromoethane in ethanol gave the ethyl 1-ethyl-3-piperidecarboxylate compound as a colorless liquid (85% yield). The ester (5 g, 27 mmol) and hydrazine hydrate (5 mL) were stirred under reflux for 4 h. Volatiles were stripped under reduced pressure. The solid was slurried in ether, filtered and dried, to give 1-ethyl piperidine-3-carbohydrazide (4.5 g, 94% yield).

[0711] (3-{5-amino-6-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl) methanol was prepared from 3-amino-6-[3-(hydroxymethyl) phenyl] pyrazine-2-carbonitrile and 1-ethyl piperidine-3-carbohydrazide in a manner analogous to that

described in the previous example: ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.15 (s, 1H); 8.07 (d, 1H); 7.7 (br s, 1H); 7.47 (t, 1H); 7.4 (t, 1H); 5.25 (t, 1H); 4.6 (d, 2H); 3.1 – 2.9 (m, 3H); 2.4 (m, 2H); 2.2 – 1.9 (m, 3H); 1.8 – 1.6 (m, 3H); 1.04 (t, 3H); MS (EI) for $\text{C}_{20}\text{H}_{25}\text{N}_7\text{O}$: 380 (MH^+).

[0712] {3-[5-amino-6-(4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol: ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.4 (m, 2H); 7.7 – 7.3 (m, 4H); 5.2 (t, 1H); 4.6 (d, 2H); MS (EI) for $\text{C}_{13}\text{H}_{12}\text{N}_6\text{O}$: 269 (MH^+).

[0713] {3-[5-amino-6-(5-methyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol: ^1H NMR (400 MHz; DMSO- d_6 + TFA): 8.8 (s, 1H); 8.1 (s, 1H); 8.05 (d, 1H); 7.45 (t, 1H); 7.4 (d, 1H); 4.6 (s, 2H); 2.5 (s, 3H); MS (EI) for $\text{C}_{19}\text{H}_{16}\text{N}_6\text{O}$: 345 (MH^+).

[0714] (3-{5-amino-6-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol: ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.1 (m, 2H); 7.6 (m, 2H); 7.45 – 7.2 (m, 6H); 5.2 (m, 1H); 4.6 (d, 2H); 4.1 (s, 2H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_6\text{O}$: 359 (MH^+).

[0715] (3-{5-amino-6-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol: ^1H NMR (400 MHz; DMSO- d_6): 8.8 (s, 1H); 8.1 (s, 1H); 7.4 (m, 3H); 5.2 (t, 1H); 4.6 (d, 2H); 1.4 (m, 9H); MS (EI) for $\text{C}_{17}\text{H}_{20}\text{N}_6\text{O}$: 325 (MH^+).

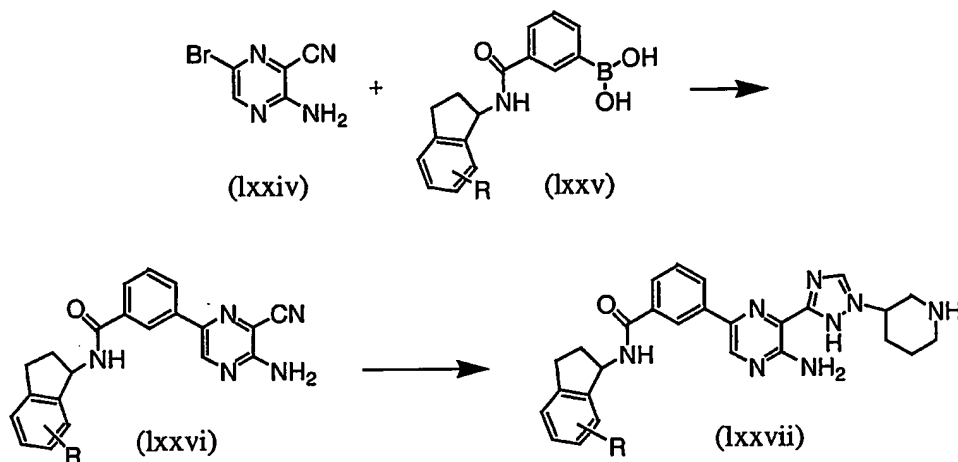
[0716] {3-[5-amino-6-(5-furan-2-yl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol: MS (EI) for $\text{C}_{17}\text{H}_{14}\text{N}_6\text{O}_2$: 335 (MH^+).

[0717] [3-(5-amino-6-{5-[3-(methyloxy)phenyl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)phenyl]methanol: MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_6\text{O}_2$: 375 (MH^+).

Example 41

[0718] Scheme 16 depicts a general synthetic route for exemplary compounds of the invention according to formula I, wherein -W-X-Y is first installed and then -A-R⁷ is installed, specifically where A is a triazole. In this example, bromo-pyrazine (lxxiv) is installed, specifically where A is a triazole. In this example, bromo-pyrazine (lxxiv) is coupled to boronic acid (lxxv) to afford intermediate (lxxvi). The cyano group of (lxxvi) is converted to, for example, a triazole to give (lxxvii), a compound according to formula I.

Scheme 16



3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide

[0719] (3-{(1S)-2,3-dihydro-1H-inden-1-ylamino}carbonyl)phenyl)boronic acid: To a suspension of 3-carboxybenzene boronic acid (7 g, 42.4 mmol) in dry methylene chloride (200 mL) were added thionyl chloride (50 mL) and 3 drops of DMF. The resulting mixture was stirred for 2 h at 50 °C. The clear solution was evaporated to dryness to give the crude acid chloride. To that residue were added (S)-aminoindane (5.64 g, 42.4 mmol) and triethylamine (12.8 mL, 3 eq) in THF (50 mL). The reaction mixture was allowed to stir overnight and then poured into water (100 mL), extracted with EtOAc (3x50 mL). The combined organic extracts were washed with brine, HCl (0.1 M, 20 mL), dried (Na₂SO₄) and concentrated to give white solid. The solid was washed with hexane to give crude title compound (6.2 g): MS (EI) for C₁₆H₁₆BNO₃: 282 (MH⁺).

[0720] To a mixture of (3-{(1S)-2,3-dihydro-1H-inden-1-ylamino}carbonyl)phenyl) boronic acid (6.2 g, 22.6 mmol) and 3-amino-6-bromopyrazine-2-carbonitrile (4.3 g, 21.8 mmol) in DMF (40 mL) was added triethylamine (9.2 mL) and Pd(dppf)₂Cl₂ (10 mol% yield). The deep black mixture was stirred overnight at 85 °C, cooled to room temperature and EtOAc (200 mL) was added. The organic layer was passed through a plug of Celite. The pale yellow solution was washed with brine (3x20 mL), HCl (1 M aqueous solution (10 mL), sodium carbonate (30 mL), dried (Na₂SO₄) and evaporated to dryness. The residue was recrystallized from hot EtOAc to give 3-(5-amino-6-cyanopyrazine-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl] benzamide (2.4 g, 30.9% yield): ¹H NMR (400 MHz;

CD₃OD): 8.9 (m, 2H); 8.45 (s, 1H); 8.15 (d, 1H); 7.92 (d, 1H); 7.65 - 7.38 (m, 3H); 7.22 (m, 4H); 5.6 (m, 3H); 3.03-2.85 (m, 2H); 2.42 (m, 1H); 2.0 (m, 1H); MS (EI) for C₂₁H₁₇N₅O: 356 (MH⁺).

[0721] A solution of 3-(5-amino-6-cyanopyrazine-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl] benzamide (2.4 g, 6.6 mmol) and 1,1-dimethyl (3S)-3-(hydrazinocarbonyl)piperidine-1-carboxylate (2.4 g, 10.1 mmol) in NMP (3 mL) was heated to 180-200 °C for 1h. The solvent was removed under reduced pressure. The residue obtained was stirred with CH₂Cl₂/TFA (20% v/v) for 30 min., concentrated and purified by hplc. The compound was converted to the hydrochloride salt using a 4 M solution of HCl in dioxane to afford 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide: ¹H NMR (400 MHz; CD₃OD): 8.65 (s, 1H); 8.6 (d, 1H); 7.6 (t, 1H); 7.7 - 7.8 (m, 4H); 5.7 (t, 1H); 3.75 - 3.4 (m, 4H); 3.8 - 3.6 (m, 3H); 3.35 - 2.8 (m, 3H); 2.6 (m, 1H); 2.4 (m, 2H); 2.1 - 2.0 (m, 4H); 1.3 (t, 3H); MS (EI) for C₂₇H₂₈N₈O: 481 (MH⁺).

[0722] 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl] benzamide: Using (1-R,2S) + cis-1-aminoindan-2-ol and 3-carboxybenzene boronic acid, the title compound was prepared according to the general scheme: ¹H NMR (400 MHz; CD₃OD): 8.86 (s, 1H); 8.6 (s, 1H); 8.5 (d, 1H); 8.4 (d, 1H); 7.94 (d, 1H); 7.63 (m, NH); 7.6 (t, 1H); 7.25 (m, 3H); 5.52 (m, 1H); 4.55 (m, 1H); 3.2 (m, 4H); 2.93 (m, 4H); 2.75 (m, 1H); 2.11 (m, 11H); 1.68 (m, 2H); 1.5 (m, 2H); MS (EI) for C₂₇H₂₈N₈O₂: 497 (MH⁺).

[0723] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0724] 3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(2-chlorophenyl)methyl]benzamide: ¹H NMR (400 MHz; DMSO-d₆): 9.3 (t, 1H); 9.0 (m, 2H); 8.8 (s, 1H); 8.73 (s, 1H); 8.45 (d, 1H); 7.8 (d, 1H); 7.75 (br s, 1H, NH); 7.6 (t, 1H); 7.49 (d, 1H); 7.41 (d, 1H); 7.3 (m, 2H); 4.6 (d, 2H); 3.6 - 2.9 (m, 4H); 2.2 (m, 1H); 1.8 (m, 4H); MS (EI) for C₂₅H₂₅N₈OCl HCl: 489 (MH⁺).

[0725] 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(2-fluorophenyl)methyl]benzamide: ¹H NMR (400 MHz; DMSO-d₆): 9.5 (m, 1H); 9.3 - 9.1 (m, 2H); 8.85 (s, 1H); 8.78 (s, 1H); 8.43 (d, 1H); 7.9 (d, 1H); 7.6 (t, 1H); 7.42 (m, 4H);

4.5 (d, 2H); 3.58 – 2.8 (m, 5H); 2.12 – 1.83 (m, 4H); MS (EI) for $C_{25}H_{25}N_8F$ HCl: 473 (MH^+).

[0726] 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(3-fluorophenyl)methyl]benzamide: 1H NMR (400 MHz; DMSO- d_6): 9.4 (m, 1H); 9.0 (m, 2H); 8.9 (s, 1); 8.7 (s, 1H); 8.4 (d, 1H); 7.9 (m, 1H); 7.8 (br s, NH); 7.6 (t, 1H); 7.2 (m, 2H); 7.1 (m, 1H); 4.6 (d, 2H); 3.9 – 2.4 (m, 5H); 2.2 – 1.8 (m, 4H); MS (EI) for $C_{25}H_{25}N_8F$ HCl: 473 (MH^+).

[0727] 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(4-fluorophenyl)methyl]benzamide: 1H NMR (400 MHz; DMSO- d_6): 9.5 (m, 1H); 9.35 – 9.1 (m, 2H); 8.9 (s, 1H); 8.8 (s, 1H); 8.4 (d, 1H); 7.9 (d, 1H); 7.6 (t, 1H); 7.4 (m, 4H); 4.5 (d, 2H); 3.6 – 2.9 (m, 5H); 2.2 (m, 1H); 1.8 (m, 3H); MS (EI) for $C_{25}H_{25}N_8OF$ HCl: 473 (MH^+).

[0728] 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(3-chlorophenyl)methyl]benzamide: 1H NMR (400 MHz; DMSO- d_6): 9.3 (t, 1H); 9.0 (m, 2H); 8.9 (s, 1H); 8.7 (s, 1H); 8.4 (d, 1H); 7.9 (d, 1H); 7.6 (t, 1H); 7.4 (m, 2H); 7.2 (m, 2H); 4.5 (d, 2H); 3.6 – 2.9 (m, 5H); 2.2 (m, 1H); 1.8 (m, 3H); MS (EI) for $C_{25}H_{25}N_8OCl$ HCl: 489 (MH^+).

[0729] 3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(4-chlorophenyl)methyl]benzamide : 1H NMR (400 MHz; DMSO- d_6): 9.3 (t, 1H); 8.9 (m, 2H); 8.67 (s, 1H); 8.4 (d, 1H); 7.8 (d, 1H); 7.7 (m, NH); 7.6 (t, 1H); 7.2 (m, 4H); 4.5 (d, 2H); 3.6 (d, 1H); 3.4 – 3.1 (m, 3H); 3.0 (m, 1H); 2.1 (m, 1H); 1.8 (m, 3H); MS (EI) for $C_{25}H_{25}N_8OCl$ HCl: 489 (MH^+).

Example 42

3-(Methylamino)-6-phenylpyrazine-2-carboxamide

[0730] Methyl 3-chloro-6-phenylpyrazine-2-carboxylate: To a mixture of acetic acid/hydrochloric acid (1:1, 20 mL) at 0° C was added methyl 3-amino-6-phenylpyrazine-2-carboxylate (1.02 g, 4.50 mmol). Aqueous solution of sodium nitrite (0.77 g, 11.2 mmol) was added drop wise. During the addition, the temperature of the reaction was controlled at 0-5° C. The reaction was slowly warmed to room temperature and stirred overnight. The reaction mixture was extracted with ethyl acetate (2 X 50 mL). The

organic layer was washed with water (30 mL), saturated aqueous sodium chloride (30 mL), dried over magnesium sulfate, filtered and concentrated at reduced pressure to afford the crude methyl 3-hydroxy-6-phenylpyrazine-2-carboxylate which was used in the next step without any purification. The crude product was treated with phosphorus oxychloride (30 mL) and heated to 90° C for 2 h. The reaction was cooled to room temperature and poured into ice-water mixture and extracted with ethyl acetate (3 X 40 mL). The organic layer was washed with water (30 mL), saturated aqueous sodium chloride (30 mL), dried over magnesium sulfate, filtered, and concentrated to give crude product. Column purification on silica (8:2 hexanes/ethyl acetate) afforded methyl 3-chloro-6-phenylpyrazine-2-carboxylate (0.22 g, 20% yield) as yellow solid: ¹H NMR (400 MHz, CDCl₃): δ 8.90 (s, 1H), 8.02 (t, 2H), 7.40 (t, 1H), 7.20 (br s, 2H), 3.90 (s, 3H); MS (EI) for C₁₂H₉N₂O₂Cl: 249 (MH⁺).

[0731] 3-(Methylamino)-6-phenylpyrazine-2-carboxamide: To a solution of methyl 3-chloro-6-phenylpyrazine-2-carboxylate (0.13 g, 0.53 mmol) in tetrahydrofuran (3 mL) was added 0.65 mL of 2.0 N methylamine in THF (1.30 mmol). The reaction was heated in a sealed high-pressure test tube to 90 -95° C overnight. The reaction was cooled to room temperature and diluted with ethyl acetate (100 mL). The organic layer was washed 2.0 N aqueous hydrochloric acid (20 mL), saturated aqueous sodium bicarbonate (30 mL), and saturated aqueous sodium chloride (30 mL). The organic layer was dried over magnesium sulfate, filtered and concentrated at reduced pressure to afford a yellowish product. Trituration with methanol (2 mL) gave a solid, which was filtered and washed with ether to afford methyl 3-(methylamino)-6-phenylpyrazine-2-carboxylate (31 mg, 24%) as yellow solid: MS (EI) for C₁₃H₁₃N₃O₂: 244 (MH⁺). This intermediate ester was directly used in the next step.

[0732] A solution of methyl 3-(methylamino)-6-phenylpyrazine-2-carboxylate (31 mg, 0.13 mmol) in methanol (5 mL) was saturated with ammonia at 0 °C. The pressure tube was sealed and heated at 80-85 °C overnight. The reaction was cooled to room temperature and the solvent removed at reduced pressure to afford crude solid product. Reverse-phase HPLC purification gave 3-(methylamino)-6-phenylpyrazine-2-carboxamide (7 mg, 25% yield) as a yellow solid: ¹H NMR (400 MHz, CDCl₃): δ 8.42 (s, 1H), 8.50 (br s, 1H), 7.90 (d, 3H), 7.45 (m, 2H), 7.40 (m, 1H), 3.10 (br s, 3H); MS (EI) for C₁₂H₁₂N₄O: 229 (MH⁺).

[0733] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0734] 6-phenyl-3-[(phenylmethyl)amino]pyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 9.01 (br t, 1H), 8.70 (s, 1H), 7.90 (m, 3H), 7.35 (m, 8H), 5.60 (br s, 1H), 4.80 (d, 2H); MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_4\text{O}$: 305 (MH^+).

[0735] 6-phenyl-3-(propylamino)pyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 8.70 (s, 1H), 8.70 (s, 1H), 8.60 (br s, 1H), 7.44 (t, 2H), 7.40 (t, 1H), 5.50 (br s, 1H), 3.50 (m, 2H), 1.70 (m, 2H), 1.02 (t, 3H); MS (EI) for $\text{C}_{14}\text{H}_{16}\text{N}_4\text{O}$: 257 (MH^+).

Example 43

3-Amino-N-methyl-6-(5-cyclopentyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide

[0736] 3-Amino-6-bromo-N-methylpyrazine-2-carboxamide (2.0 g, 8.7 mmol) and copper (I) cyanide (3.9 g, 43.5 mmol) were taken into DMF (20 mL) and the mixture was heated to reflux for two hours. The mixture was allowed to cool to room temperature then diluted with ethyl acetate (250 mL). The insoluble residue was removed by filtration and the filtrate partitioned with water. The organic layer was separated and washed with water (2x) then brine and dried over anhydrous magnesium sulfate. Filtration and concentration afforded 3-amino-6-cyano-N-methylpyrazine-2-carboxamide (500 mg, 33% yield) as a white solid: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): 8.84 (br m, 1H), 8.67 (br s, 1H), 8.65 (s, 1H), 2.77 (d, 3H).

[0737] 3-Amino-6-cyano-N-methylpyrazine-2-carboxamide (839 mg, 4.7 mmol) was suspended in ethanol (25 mL) followed by addition of 50% aqueous hydroxylamine (2 mL, 31.3 mmol) and the mixture was stirred at room temperature for one hour. The suspension obtained was diluted with water (25 mL) and the insoluble solid collected by filtration and washed with water. The solid was dried *in vacuo* to give 3-amino-6-[(hydroxyamino)(imino)methyl]-N-methylpyrazine-2-carboxamide (664 mg, 67% yield) as a white solid: MS (EI) for $\text{C}_7\text{H}_{10}\text{N}_6\text{O}_2$: 211 (MH^+).

[0738] 3-Amino-6-[(hydroxyamino)(imino)methyl]-N-methylpyrazine-2-carboxamide (40 mg, 0.19 mmol) was taken into DMF (1.0 mL) followed by addition of PL-DIPAM (diisopropylaminomethyl) resin (200 mg, 2.33 mmol/g), THF (1.0 mL) and cyclopentanecarbonyl chloride (23.1 μL , 0.19 mmol). The mixture was agitated for one

hour then filtered and the resin rinsed with THF (1.0 mL). To the filtrate solution was added tetra-*N*-butylammonium fluoride (TBAF) (1.0 M in THF, 20 0uL) and the solution was allowed to stir at room temperature over 12 hours. The mixture was concentrated and the residue partitioned with ethyl acetate and water. The organic layer was washed with water (1x), 1.0 M aqueous hydrochloric acid (1x) then brine and dried over anhydrous magnesium sulfate. The material was filtered and concentrated then the residue purified by silica gel flash chromatography using ethyl acetate:hexanes (1:1) as eluent. The pure fractions were combined, concentrated and dried in vacuo to give 3-amino-*N*-methyl-6-(5-cyclopentyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide (23.3 mg, 43% yield) as a colorless amorphous residue: MS (EI) for $C_{13}H_{16}N_6O_2$: 289 (MH^+).

[0739] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0740] 3-Amino-*N*-methyl-6-(5-pyridin-3-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide: MS (EI) for $C_{13}H_{11}N_7O_2$: 298 (MH^+).

[0741] 3-Amino-*N*-methyl-6-(5-cyclohexyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide: MS (EI) for $C_{14}H_{18}N_6O_2$: 303 (MH^+).

[0742] 3-Amino-*N*-methyl-6-(5-phenyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide: MS (EI) for $C_{14}H_{12}N_6O_2$: 297 (MH^+).

[0743] 3-Amino-*N*-methyl-6-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide: MS (EI) for $C_{15}H_{14}N_6O_2$: 311 (MH^+).

[0744] 3-Amino-*N*-methyl-6-(5-methyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide: MS (EI) for $C_9H_{10}N_6O_2$: 235 (MH^+).

[0745] 3-Amino-*N*-methyl-6-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide: MS (EI) for $C_{16}H_{16}N_6O_2$: 325 (MH^+).

[0746] 3-Amino-*N*-methyl-6-(5-pyridin-4-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide: MS (EI) for $C_{13}H_{11}N_7O_2$: 298 (MH^+).

[0747] 3-Amino-*N*-methyl-6-[5-(1,1-dimethylethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide: MS (EI) for $C_{12}H_{16}N_6O_2$: 276 (MH^+).

[0748] 3-Amino-*N*-methyl-6-(5-furan-2-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide: MS (EI) for $C_{12}H_{10}N_6O_3$: 286 (MH^+).

Example 44

3-Amino-N-methyl-6-[3-(5-phenyl-1,2,4-oxadiazol-3-yl)phenyl]pyrazine-2-carboxamide

[0749] 3-Amino-6-(3-cyanophenyl)-N-methylpyrazine-2-carboxamide (155 mg, 0.61mmol) was suspended in ethanol (2 mL) followed by addition of 50% aqueous hydroxylamine (250 uL, 4.1 mmol) and the mixture was heated to reflux for 30 minutes then allowed to cool to room temperature. The suspension was diluted with water (2 mL) and the solid product collected by filtration and dried *in vacuo* to give crude 3-amino-N-methyl-6-{3-[(hydroxyamino)(imino)methyl]phenyl}pyrazine-2-carboxamide (125 mg, 72% yield) as a yellow solid which was carried forward without further purification: MS (EI) for C₁₃H₁₅N₆O₂: 287 (MH⁺).

[0750] 3-Amino-N-methyl-6-{3-[(hydroxyamino)(imino)methyl]phenyl}pyrazine-2-carboxamide (40mg, 0.14mmol) was suspended in DMF (750 uL) followed by addition of PL-DIPAM (diisopropylaminomethyl) resin (100 mg, 2.33 mmol/g) and benzoyl chloride (20 uL, 0.17 mmol). The mixture was agitated for one hour then diluted with ethyl acetate and the spent resin removed by filtration. The organic filtrate was washed with water (3x), brine then dried over anhydrous magnesium sulfate. The mixture was then filtered and concentrated and the solid residue suspended in THF (1 mL) followed by addition of tetra-N-butylammonium fluoride (TBAF) (1.0 M in THF, 100 uL) and the solution was allowed to stir at room temperature over 12 hours. The mixture was concentrated and the residue partitioned with ethyl acetate and water. The organic layer was washed with water (1x), 1.0 M aqueous hydrochloric acid (1x) then brine and dried over anhydrous magnesium sulfate. The material was filtered and concentrated then the residue purified by preparative reverse phase HPLC (water/acetonitrile/0.1% TFA eluent). The pure fractions were combined and lyophilized to afford 3-amino-N-methyl-6-[3-(5-phenyl-1,2,4-oxadiazol-3-yl)phenyl]pyrazine-2-carboxamide (10.4 mg, 20% yield) as a yellow solid: MS (EI) for C₂₀H₁₇N₆O₂: 372 (MH⁺).

[0751] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:

[0752] 3-Amino-N-methyl-6-{3-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]phenyl}pyrazine-2-carboxamide: MS (EI) for C₂₁H₁₉N₆O₂: 386 (MH⁺).

[0753] 3-Amino-N-methyl-6-{3-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]phenyl}pyrazine-2-carboxamide: MS (EI) for C₂₁H₁₉N₆O₂: 400 (MH⁺).

Example 45**3-Amino-N-methyl-6-{3-[(2-phenylethyl)oxy]phenyl}pyrazine-2-carboxamide**

[0754] 3-Amino-6-bromo-N-methylpyrazine-2-carboxamide (1.34 g, 5.8 mmol) and 3-hydroxyphenylboronic acid (1.2 g, 8.7 mmol) were taken into DMF (10 mL) followed by addition of [1,1'-bis(diphenylphosphino)ferrocene]dichloropalladium (II) dichloromethane complex (440 mg, 0.54 mmol) and triethylamine (4.8 mL, 35 mmol). The mixture was heated to 95 °C for 12 hours then cooled to room temperature and diluted with an excess of ethyl acetate. The organic solution was washed with 0.5 M aqueous hydrochloric acid (1x), water (3x) then brine and dried over anhydrous magnesium sulfate. The mixture was then filtered and the organic solution concentrated and the residue purified by silica gel flash chromatography using ethyl acetate:hexanes (3:1) eluent. The pure fractions were combined and concentrated then dried *in vacuo* to give 3-amino-6-(3-hydroxyphenyl)-N-methylpyrazine-2-carboxamide (1.01 g, 71% yield) as a yellow solid: MS (EI) for C₁₂H₁₃N₄O₂: 245 (MH⁺).

[0755] 3-Amino-6-(3-hydroxyphenyl)-N-methylpyrazine-2-carboxamide (179 mg, 0.8 mmol) was taken into DMF (3 mL) followed by addition of cesium carbonate (1.3 g, 4 mmol) and 2-(bromoethyl)benzene (550 uL, 4 mmol). The mixture was heated to 85 °C for 12 hours then allowed to cool to room temperature. The mixture was diluted with an excess of ethyl acetate and washed with water (3x) then brine and dried over anhydrous magnesium sulfate. Filtration and concentration *in vacuo* followed by purification of the residue using silica gel flash chromatography using ethyl acetate:hexanes (2:3) as eluent. The pure fractions were combined and concentrated then dried *in vacuo* to give 3-amino-N-methyl-6-{3-[(2-phenylethyl)oxy]phenyl}pyrazine-2-carboxamide (86 mg, 31% yield) as a yellow solid. ¹H NMR (400 MHz, d₆-DMSO): 8.83 (s, 1H), 8.82 (br m, 1H), 7.74 (d, 1H), 7.67 (tr, 1H), 7.38-7.31 (m, 6H), 7.26-7.22 (m, 1H), 6.95 (d tr, 1H), 4.31 (tr, 2H), 3.08 (tr, 2H), 2.85 (d, 3H); MS (EI) for C₂₀H₂₁N₄O₂: 349 (MH⁺).

Example 46**3-Amino-N-methyl-6-(3-{[2-(4-methylpiperidin-1-yl)ethyl]oxy}phenyl)pyrazine-2-carboxamide**

- [0756] 3-Amino-6-(3-hydroxyphenyl)-*N*-methylpyrazine-2-carboxamide (1.01 g, 4.1 mmol) was taken into DMF (10 mL) followed by addition of 1,2-dibromoethane (10 mL) and cesium carbonate (2.7 g, 8.2 mmol). The mixture was warmed to 85 °C over 5 days followed by addition of another aliquot of cesium carbonate and 1,2-dibromoethane and heating was continued an additional 4 hours. The mixture was diluted with an excess of ethyl acetate and washed with water (3x) then brine and dried over anhydrous magnesium sulfate. Filtration and concentration *in vacuo* followed by purification of the residue using silica gel flash chromatography using ethyl acetate:hexanes (1:1) as eluent. The pure fractions were combined and concentrated then dried *in vacuo* to give 3-amino-6-{3-[(2-bromoethyl)oxy]phenyl}-*N*-methylpyrazine-2-carboxamide (486 mg, 34% yield) as a yellow solid: MS (EI) for C₁₄H₁₆N₄O₂Br: 351, 353 (MH⁺).
- [0757] 3-Amino-6-{3-[(2-bromoethyl)oxy]phenyl}-*N*-methylpyrazine-2-carboxamide (25 mg, 0.07 mmol) was taken into acetonitrile (1 mL) followed by addition of 4-methylpiperidine (58 mg, 0.58 mmol) and the mixture was stirred at room temperature over 12 hours. The crude reaction mixture was purified directly by preparative reverse phase HPLC (water/acetonitrile/0.1% TFA eluent). The pure fractions were combined and lyophilized to afford 3-amino-*N*-methyl-6-(3-{[2-(4-methylpiperidin-1-yl)ethyl]oxy}phenyl)pyrazine-2-carboxamide (20.2 mg, 60% yield) as an amorphous solid: MS (EI) for C₂₀H₂₈N₅O₂: 371 (MH⁺).
- [0758] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following compounds of the invention were prepared:
- [0759] 3-Amino-6-[3-({2-[4-(2-hydroxyethyl)piperidin-1-yl]ethyl}oxy)phenyl]-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₂₁H₃₀N₅O₃: 401 (MH⁺).
- [0760] 3-Amino-6-(3-{[2-(3-hydroxypyrrolidin-1-yl)ethyl]oxy}phenyl)-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₁₈H₂₄N₅O₃: 359 (MH⁺).
- [0761] 3-Amino-*N*-methyl-6-[3-({2-[4-(phenylmethyl)piperidin-1-yl]ethyl}oxy)phenyl]pyrazine-2-carboxamide: MS (EI) for C₂₆H₃₂N₅O₂: 447 (MH⁺).
- [0762] 3-Amino-*N*-methyl-6-{3-[(2-piperidin-1-ylethyl)oxy]phenyl}pyrazine-2-carboxamide: MS (EI) for C₁₉H₂₆N₅O₂: 357 (MH⁺).
- [0763] 3-Amino-6-[3-({2-[ethyl(phenylmethyl)amino]ethyl}oxy)phenyl]-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₂₃H₂₈N₅O₂: 407 (MH⁺).

- [0764] 3-Amino-6-(3-{[2-(diethylamino)ethyl]oxy}phenyl)-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₁₈H₂₆N₅O₂: 344 (MH⁺).
- [0765] 3-Amino-*N*-methyl-6-(3-{[2-(4-phenylpiperazin-1-yl)ethyl]oxy}phenyl)pyrazine-2-carboxamide: MS (EI) for C₂₄H₃₀N₆O₂: 434 (MH⁺).
- [0766] 3-Amino-6-[3-({2-[methyl(phenylmethyl)amino]ethyl}oxy)phenyl]-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₂₂H₂₆N₅O₂: 393 (MH⁺).
- [0767] 3-Amino-*N*-methyl-6-[3-({2-(morpholin-1-ylethyl)oxy}phenyl)pyrazine-2-carboxamide: MS (EI) for C₁₈H₂₄N₅O₃: 359 (MH⁺).
- [0768] 3-Amino-6-[3-({2-[(cyclopropyl)(propyl)amino]ethyl}oxy)phenyl]-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₂₁H₃₀N₅O₂: 384 (MH⁺).
- [0769] 3-Amino-*N*-methyl-6-(3-{[2-(4-ethylpiperazin-1-yl)ethyl]oxy}phenyl)pyrazine-2-carboxamide: MS (EI) for C₂₀H₂₉N₆O₂: 385 (MH⁺).
- [0770] 3-Amino-*N*-methyl-6-[3-({2-(pyrrolidin-1-ylethyl)oxy}phenyl)pyrazine-2-carboxamide: MS (EI) for C₁₈H₂₄N₅O₂: 343 (MH⁺).
- [0771] 3-Amino-*N*-methyl-6-(3-{[2-(4-methylpiperazin-1-yl)ethyl]oxy}phenyl)pyrazine-2-carboxamide: MS (EI) for C₁₉H₂₇N₆O₂: 371 (MH⁺).
- [0772] 3-Amino-6-[3-({2-[4-(2-hydroxyethyl)piperazin-1-yl]ethyl}oxy)phenyl]-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₂₀H₂₉N₆O₃: 402 (MH⁺).
- [0773] 3-Amino-6-(3-{[2-(3,4-dihydroisoquinolin-2(1*H*)-yl)ethyl]oxy}phenyl)-*N*-methylpyrazine-2-carboxamide: MS (EI) for C₂₃H₂₆N₅O₂: 405 (MH⁺).
- [0774] Ethyl 4-{2-[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)oxy]ethyl} piperazine-1-carboxylate: MS (EI) for C₂₁H₂₉N₆O₄: 430 (MH⁺).

Example 47

3-Amino-*N*-methyl-6-(3-{[(phenylmethyl)thio]methyl}phenyl)pyrazine-2-carboxamide

- [0775] 3-Amino-6-(3-hydroxymethylphenyl)-*N*-methylpyrazine-2-carboxamide (130 mg, 0.5 mmol) was suspended in THF (5 mL) followed by addition of pyridine (200 μ L, 2.5 mmol) and the mixture was cooled to -78 °C. Thionyl chloride (100 μ L, 1.4 mmol) was added by syringe and the mixture was allowed to slowly warm to room temperature. An additional aliquot of thionyl chloride was added at room temperature and the mixture was

allowed to stir an additional hour. The solvent was removed and the residue partitioned with ethyl acetate and 0.1 M aqueous hydrochloric acid. The organic layer was washed with saturated aqueous sodium bicarbonate then brine and dried over anhydrous magnesium sulfate. The mixture was filtered, concentrated and dried *in vacuo* to afford the crude benzyl chloride, which was taken on without further purification.

[0776] The above chloride was taken into acetonitrile (3 mL) followed by addition of benzyl mercaptan (100 μ L, 0.86 mmol) then cesium carbonate (350 mg, 1.1 mmol) and the mixture was stirred at room temperature over one hour. The solvent was removed *in vacuo* and the residue partitioned with ethyl acetate and water. The organic layer was washed with brine and dried over anhydrous magnesium sulfate. The mixture was filtered, concentrated then dried *in vacuo* and the residue purified by silica gel flash chromatography using ethyl acetate:hexanes (1:1) as eluent. The pure fractions were combined, concentrated and dried *in vacuo* to give 3-amino-N-methyl-6-(3-[[[(phenylmethyl)thio]methyl]phenyl]pyrazine-2-carboxamide (77.1 mg, 40% yield) as a yellow amorphous solid: ^1H NMR (400 MHz, CDCl_3): 8.61 (s, 1H), 7.98 (br s, 1H), 7.76-7.74 (m, 2H), 7.42 (tr, 1H), 7.35-7.24 (m, 8H), 3.68 (s, 2H), 3.65 (s, 2H), 3.05 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{21}\text{N}_4\text{O}_2$: 366 (MH^+).

Example 48

3-Amino-N-methyl-6-(3-[[[(phenylmethyl)sulfonyl]methyl]phenyl]pyrazine-2-carboxamide

[0777] 3-Amino-N-methyl-6-(3-[[[(phenylmethyl)thio]methyl]phenyl]pyrazine-2-carboxamide (73 mg, 0.2 mmol) was taken into dichloromethane (3 mL) followed by addition of *m*-CPBA (99 mg, 0.4 mmol) and the mixture was stirred for 30 minutes at room temperature. The solvent was removed and the residue partitioned with ethyl acetate and saturated aqueous sodium bicarbonate. The organic layer was washed with brine and dried over anhydrous magnesium sulfate. The mixture was filtered, concentrated and the residue suspended in a minimum of hot methanol to give a yellow solid. The material was collected by filtration and further purified by preparative reverse phase HPLC (water/acetonitrile/0.1% TFA eluent). The pure fractions were combined and lyophilized to afford 3-amino-N-methyl-6-(3-[[[(phenylmethyl)sulfonyl]methyl]

phenyl)pyrazine-2-carboxamide (35mg, 44% yield) as a yellow solid: MS (EI) for $C_{20}H_{21}N_4O_3S$: 398 (MH^+).

Example 49

3-Amino-N-methyl-6-(3-([(phenylmethyl)amino]methyl)phenyl)pyrazine-2-carboxamide

[0778] Methyl 3-amino-6-(3-chloromethylphenyl)-*N*-methylpyrazine-2-carboxylate (30.6 mg, 0.11 mmol.) was taken into acetonitrile (2 mL) and THF (1 mL) followed by addition of benzylamine (12 μ L, 0.11 mmol) then diisopropylethylamine (20 μ L, 0.11 mmol) and the mixture was allowed to stir at room temperature over six days. The mixture was then concentrated *in vacuo* and the residue purified by silica gel flash chromatography initially using ethyl acetate as eluent followed by elution with 10% methanol in dichloromethane. The pure fractions were combined and concentrated then dried *in vacuo* to give methyl 3-amino-6-(3-([(phenylmethyl)amino]methyl)phenyl)pyrazine-2-carboxylate (4.9 mg, 13% yield) as a yellow amorphous solid: MS (EI) for $C_{20}H_{21}N_4O_2$: 349 (MH^+).

[0779] Methyl 3-amino-6-(3-([(phenylmethyl)amino]methyl)phenyl)pyrazine-2-carboxylate (4.9 mg, 0.014 mmol) was taken into methanol (2 mL) and the solution was saturated with methylamine gas. The mixture was allowed to stir for two hours in a sealed vessel then concentrated and dried *in vacuo* to give 3-amino-N-methyl-6-(3-([(phenylmethyl)amino]methyl)phenyl)pyrazine-2-carboxamide (4.9 mg, 100% yield) as a yellow amorphous solid: MS (EI) for $C_{20}H_{22}N_5O$: 348 (MH^+).

Example 50

3-Amino-N-methyl-6-(3-([(phenylmethyl)oxy]methyl)phenyl)pyrazine-2-carboxamide

[0780] Methyl 3-amino-6-(3-hydroxymethylphenyl)-*N*-methylpyrazine-2-carboxylate (75 mg, 0.3 mmol) was taken into THF (2 mL) followed by addition of pyridine (120 μ L, 1.5 mmol) and the solution was cooled to 0 °C. Thionyl chloride (42 μ L, 0.6 mmol) was added to the mixture by syringe followed by warming to room temperature. The mixture was then stirred for two hours and partitioned with ethyl acetate and 1 M aqueous hydrochloric acid. The organic layer was washed with water (1x) then brine and dried over anhydrous sodium sulfate. The mixture was filtered and concentrated followed by silica gel flash chromatography using ethyl acetate:hexanes (1:1) as eluent. The pure

fractions were combined and concentrated then dried *in vacuo* to give methyl 3-amino-6-(3-chloromethylphenyl)-*N*-methylpyrazine-2-carboxylate (30 mg, 37% yield) as a yellow solid: MS (EI) for $C_{13}H_{13}N_3O_2Cl$: 278, 280 (MH^+).

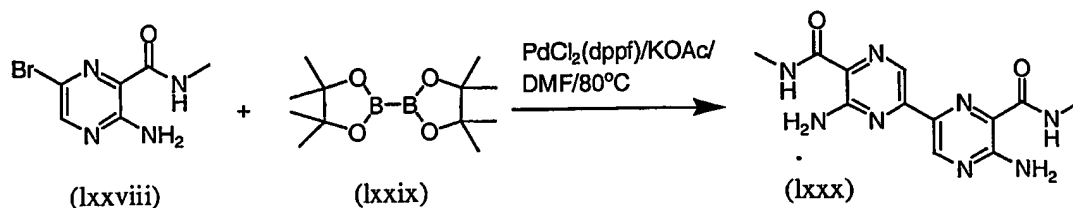
[0781] Methyl 3-amino-6-(3-chloromethylphenyl)-*N*-methylpyrazine-2-carboxylate (18 mg, 0.06 mmol) and silver oxide (15 mg, 0.06 mmol) were added to benzyl alcohol (2 mL) and the mixture was heated to 110 °C over 15 hours. The benzyl alcohol was removed *in vacuo* and the residue purified by silica gel flash chromatography using ethyl acetate:hexanes (1:1) as eluent. The pure fractions were combined and concentrated then dried *in vacuo* to give methyl 3-amino-6-(3-[[[(phenylmethyl)oxy]methyl]phenyl]pyrazine-2-carboxylate (5.8 mg, 28% yield) as a yellow solid: MS (EI) for $C_{20}H_{20}N_3O_3$: 350 (MH^+).

[0782] Methyl 3-amino-6-(3-[[[(phenylmethyl)oxy]methyl]phenyl]pyrazine-2-carboxylate (5.8 mg, 0.017 mmol) was taken into methanol (2 mL) and the solution was saturated with methylamine gas. The mixture was allowed to stir for two hours in a sealed vessel then concentrated. The residue was purified by silica gel flash chromatography using ethyl acetate:hexanes (2:3) as eluent. The pure fractions were combined and concentrated then dried *in vacuo* to give 3-amino-*N*-methyl-6-(3-[[[(phenylmethyl)oxy]methyl]phenyl]pyrazine-2-carboxamide (4.1 mg, 69% yield) as a yellow amorphous solid: MS (EI) for $C_{20}H_{21}N_4O_2$: 349 (MH^+).

Example 51

[0783] Scheme 17 shows that bromo-aromatic compounds were dimerized in some cases to make compounds according to formula I. In this example, bromo-pyrazine (lxxviii) is dimerized using boronic ester (lxxix), for example, to give (lxxx).

Scheme 17



5',6'-Diamino-*N,N'*-dimethyl-2,2'-bipyrazine-5,6'-dicarboxamide

[0784] 3-Amino-6-bromo-N-methylpyrazine-2-carboxamide: (50.0 mg, 0.22 mmol) was dissolved in DMF followed by addition of bis(pinacolate)diborane (1.57g, 0.43 mmol), KOAc (32.0 mg, 0.33 mmol) and PdCl₂(dppf) (10 mol%, 15.8 mg). This mixture was stirred at 80 °C under nitrogen for 12 hours. The reaction mixture was poured into water (10 mL), and extracted with ethyl acetate (3 × 20 mL). The organic layers were washed with brine (50 mL) then dried over anhydrous sodium sulfate. Filtration, concentration and purification by MPLC gave a solid that was dried to afford 8.45 mg (11.3% yield) of product: ¹H NMR (400 MHz, CDCl₃): δ 8.20 (s, 1H), 7.80 (s, 1H), 7.45 (s, 1H), 3.00 (d, 6H); MS (EI) for C₁₂H₁₄N₂O₂: 303.45 (MH⁺).

Example 52

3-Amino-6-[3-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide

[0785] 3-Amino-6-bromo-N-methylpyrazine-2-carboxamide (2.00g, 8.65 mmol) was dissolved in DMF followed by addition of 3-hydroxymethyl phenyl boronic acid (1.57g, 10.32 mmol), K₂CO₃ (3.0g, 21.7 mmol) and Pd(PPh₃)₄ (10 mol%, 1.0g). This mixture was stirred at 90 °C under nitrogen for 12 hours. The reaction mixture was poured into water (150 mL), and extracted with ethyl acetate (3 × 50 mL). The organic layers were washed with brine (50 mL) then dried over anhydrous sodium sulfate. Filtration, concentration and column chromatography on silica (50% ethylacetate/hexanes) gave a solid which was dried to afforded 2.02 g (92% yield) of the title compound: ¹H NMR (400 MHz, d₆-DMSO): δ 8.80 (s, 1H), 8.74 (br, 1H), 8.01(s, 2H), 7.45 (m, 4H), 5.21(t, 1H), 4.60 (d, 2H), 2.81(d, 3H); MS (EI) for C₁₃H₁₄N₄O₂: 259.38 (MH⁺).

[0786] 3-Amino-6-[3-(bromomethyl)phenyl]-N-methylpyrazine-2-carboxamide 3-Amino-6-[3-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide (1.00g, 3.87 mmol) was dissolved in dichloromethane followed by addition of triphenylphosphine (1.83g, 6.98 mmol) and carbon tetrabromide (2.13g, 6.42 mmol). This mixture was stirred under nitrogen overnight. Concentration and column chromatography on silica (20% → 50% ethylacetate/hexanes) gave a solid which was dried to afforded 0.42 g (39 % yield) of intermediate 2. ¹H NMR (400 MHz, CDCl₃): δ 8.60 (s, 1H), 7.45 (m, 5H), 4.60 (d, 2H), 3.05(d, 3H); MS (EI) for C₁₃H₁₃BrN₄O: 323.20 (M+2).

[0787] 3-Amino-6-(3-{[5-methyl-pyrazine-2-ylmethyl]-amino}-methyl)-phenyl)-pyrazine-2-carboxylic acid methylamide: Intermediate 2 (31.0 mg, 1.12 mmol) was

dissolved into acetonitrile (0.5 M solution) followed by addition of DIPEA (1.344 mmol) and C-(5-methyl-pyrazin-2-yl)-methylamine (28.0 mg, 2.27 mmol) and was stirred overnight at RT. This reaction was poured into water and brine mixture and extracted with ethylacetate (3×20 mL). The combined organic layer was dried over anhydrous sodium sulfate. Filtration, concentration and purified by MPLC and dried to give a yellow powder. ¹H NMR (400 MHz, d₆-DMSO): δ 8.80 (s, 1H), 8.45 (s, 2H), 8.10 (s, 1H), 7.98 (m, 1H), 7.40 (m, 1H), 7.38 (m, 1H), 3.98 (s, 2H), 3.96 (s, 2H), 3.00 (s, 3H), 2.50 (s, 3H); MS (EI) for C₁₉H₂₁N₇O: 364.47 (MH⁺).

[0788] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.

[0789] 3-amino-6-(3-(((biphenyl-4-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 8.90 (m, 2H), 8.00 (m, 2H), 7.60 (m, 4H), 7.40 (m, 8H), 3.80 (d, 4H), 2.80 (d, 3H); MS (EI) for C₂₆H₂₅NO₅: 424.21 (MH⁺).

[0790] 3-amino-6-(3-(((biphenyl-2-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₄-MeOH): δ 8.60 (s, 1H), 7.80 (m, 2H), 7.35 (m, 11H), 3.75 (s, 2H), 3.65 (s, 2H), 3.30 (d, 3H); MS (EI) for C₂₆H₂₅NO₅: 424.48 (MH⁺).

[0791] 3-amino-6-(3-(((3,4-dihydro-2H-1,4-benzoxazin-6-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 8.90 (m, 2H), 8.03 (m, 2H), 7.40 (m, 2H), 6.60 (m, 3H), 5.70 (s, 1H), 4.11 (m, 2H), 3.80 (s, 2H), 3.60 (s, 2H), 3.20 (s, 2H), 2.80 (d, 3H); MS (EI) for C₂₂H₂₄N₆O₂: 405.39 (MH⁺).

[0792] 3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 9.40 (br, 2H), 8.80 (m, 2H), 8.21 (m, 2H), 7.40 (m, 4H), 4.38 (d, 4H), 2.83 (d, 3H); MS (EI) for C₂₀H₁₉NO₅F₂: 384.43 (MH⁺).

[0793] 3-amino-6-[3-(((2,5-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 9.40 (br, 2H), 8.80 (m, 2H), 8.21 (m, 2H), 7.40 (m, 4H), 4.38 (d, 4H), 2.83 (d, 3H); MS (EI) for C₂₀H₁₉NO₅F₂: 384.41 (MH⁺).

- [0794] 3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.05 (m, 2H), 7.35 (m, 6H), 3.80 (s, 4H), 2.80 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{NO}_5\text{F}_2$: 384.41 (MH^+).
- [0795] 3-amino-N-methyl-6-[3-(((4-(methylsulfonyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.05 (m, 2H), 7.80 (m, 2H), 7.60 (m, 3H), 7.40 (m, 2H), 3.80 (d, 4H), 3.20 (s, 3H), 2.80 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{23}\text{N}_5\text{O}_3\text{S}$: 426.42 (MH^+).
- [0796] 3-amino-6-(3-(((2,2-diphenylethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.05 (m, 2H), 7.60 (m, 2H), 7.22 (m, 11H), 4.20 (m, 1H), 3.80 (s, 4H), 2.80 (d, 3H); MS (EI) for $\text{C}_{27}\text{H}_{27}\text{NO}_5$: 438.48 (MH^+).
- [0797] 3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (d, 2H), 8.11 (d, 2H), 7.90 (m, 2H), 7.60 (m, 1H), 7.40 (m, 3H), 7.00 (m, 4H), 3.80 (d, 4H), 2.92 (d, 3H); MS (EI) for $\text{C}_{26}\text{H}_{25}\text{N}_5\text{O}_2$: 440.47 (MH^+).
- [0798] 3-amino-6-[3-(((4-aminophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.11 (d, 2H), 7.40 (m, 2H), 7.00 (m, 1H), 6.50 (m, 2H), 4.95 (s, 2H), 3.80 (s, 2H), 3.60 (s, 2H), 2.80 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{22}\text{N}_6\text{O}$: 363.44 (MH^+).
- [0799] 3-amino-N-methyl-6-[3-(((4-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.65 (s, 2H), 8.11 (s, 1H), 7.96 (d, 1H), 7.40 (m, 5H), 6.92 (m, 6H), 3.80 (d, 4H), 2.92 (d, 3H); MS (EI) for $\text{C}_{26}\text{H}_{25}\text{N}_5\text{O}_2$: 440.47 (MH^+).
- [0800] 3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.11 (s, 1H), 7.90 (d, 1H), 7.40 (m, 1H), 7.20 (m, 2H), 6.60 (d, 2H), 3.85 (d, 4H), 3.80 (s, 6H), 2.95 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{25}\text{N}_5\text{O}_3$: 408.47 (MH^+).
- [0801] 3-amino-6-[3-(((2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.11 (s,

1H), 7.90 (d, 1H), 7.40 (m, 6H), 3.85 (d, 4H), 2.95 (d, 3H); MS (EI) for $C_{20}H_{20}N_5OF$: 366.45 (MH^+).

[0802] 3-amino-6-[3-([(2,6-difluorophenyl)methyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d6-DMSO): δ 9.80 (s, 2H), 8.90 (m, 2H), 8.40 (s, 1H), 8.20 (s, 1H), 7.50 (m, 2H), 7.20 (m, 2H), 4.30 (d, 4H), 2.80 (d, 3H); MS (EI) for $C_{20}H_{19}N_5OF_2$: 384.42 (MH^+).

[0803] 3-amino-6-(3-([(2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)methyl]phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.40 (m, 3H), 6.80 (m, 4H), 4.20 (s, 4H), 3.80 (s, 2H), 3.60 (s, 2H), 3.00 (d, 3H); MS (EI) for $C_{22}H_{23}N_5O_3$: 406.49 (MH^+).

[0804] 3-amino-N-methyl-6-[3-([(2-piperidin-1-yl]phenyl)methyl]amino)methyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.40 (m, 6H), 3.80 (d, 4H), 2.99 (d, 3H), 2.80 (d, 4H), 1.60 (m, 6H); MS (EI) for $C_{25}H_{30}N_6O$: 431.50 (MH^+).

[0805] 3-amino-6-[3-([(2,5-dichlorophenyl)methyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d6-DMSO): δ 8.90 (m, 2H), 8.10 (m, 2H), 7.40 (m, 6H), 3.80 (d, 4H), 2.80 (d, 3H); MS (EI) for $C_{20}H_{19}N_5OCl_2$: 416.29 (M^+).

[0806] 3-amino-6-[3-([(3,5-dichlorophenyl)methyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d6-DMSO): δ 8.90 (m, 2H), 8.10 (m, 2H), 7.40 (m, 6H), 3.80 (d, 4H), 2.80 (d, 3H); MS (EI) for $C_{20}H_{19}N_5OCl_2$: 416.32 (M^+).

[0807] 3-amino-6-[3-[(2,3-dihydro-1H-inden-2-ylamino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.40 (m, 2H), 7.10 (m, 4H), 3.90 (s, 2H), 3.60 (m, 1H), 3.20 (m, 2H), 3.00 (s, 3H), 2.80 (m, 2H); MS (EI) for $C_{22}H_{23}N_5O$: 374.42 (MH^+).

[0808] 3-amino-6-[3-(1,3-dihydro-2H-isoindol-2-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d6-DMSO): δ 8.80 (m, 2H), 8.10 (m, 2H), 7.40 (m, 2H), 7.20 (m, 4H), 3.98 (s, 2H), 3.82 (s, 4H), 2.80 (d, 3H); MS (EI) for $C_{21}H_{21}N_5O$: 360.45 (MH^+).

- [0809] 3-amino-6-(3-([(1R)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.40 (m, 3H), 7.10 (m, 3H), 4.40 (m, 1H), 3.97 (m, 2H), 3.10 (m, 1H), 3.00 (s, 3H), 2.94 (m, 1H), 2.40 (m, 1H), 2.00 (m, 1H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}$: 374.45 (MH^+).
- [0810] 3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.40 (m, 3H), 7.10 (m, 3H), 4.40 (m, 1H), 3.97 (m, 2H), 3.10 (m, 1H), 3.00 (s, 3H), 2.94 (m, 1H), 2.40 (m, 1H), 2.00 (m, 1H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}$: 374.46 (MH^+).
- [0811] 3-amino-N-methyl-6-[3-([(3,4,5-trifluorophenyl)methyl]amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.40 (m, 1H), 7.30 (m, 1H), 7.08 (m, 2H), 4.40 (m, 1H), 3.80 (d, 4H), 3.00 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{OF}_3$: 402.40 (MH^+).
- [0812] 3-amino-N-methyl-6-[3-([(1R)-1-phenylethyl]amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.70 (s, 1H), 8.00 (s, 1H), 7.90 (d, 1H), 7.40 (m, 6H), 3.80 (m, 1H), 3.60 (m, 2H), 3.00 (s, 3H), 1.40 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{23}\text{N}_5\text{O}$: 362.44 (MH^+).
- [0813] 3-amino-6-[3-([(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 8.90 (m, 2H), 8.20 (s, 1H), 8.02 (d, 1H), 7.50 (m, 4H), 7.20 (m, 3H), 4.80 (br, 1H), 4.40 (m, 1H), 4.00 (m, 3H), 2.95 (m, 5H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_2$: 390.28 (MH^+).
- [0814] 3-amino-N-methyl-6-[3-([(4-(2-thienyl)phenyl)methyl]amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.90 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.60 (m, 2H), 7.40 (m, 6H), 7.10 (m, 1H), 3.80 (m, 4H), 3.00 (s, 3H); MS (EI) for $\text{C}_{24}\text{H}_{23}\text{N}_5\text{OS}$: 430.42 (MH^+).
- [0815] 3-amino-N-methyl-6-[3-([(3-(1,2,3-thiadiazol-4-yl)phenyl)methyl]amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 9.60 (s, 1H), 8.80 (m, 2H), 8.15 (m, 4H), 7.40 (m, 5H), 4.00 (s, 1H), 3.80 (s, 3H), 2.80 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_7\text{OS}$: 432.42 (MH^+).

- [0816] 3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (m, 1H), 7.40 (m, 5H), 3.80 (d, 4H), 2.95 (s, 3H); MS (EI) for $\text{C}_{20}\text{H}_{19}\text{N}_5\text{OFCl}$: 400.41 (MH^+).
- [0817] 3-amino-N-methyl-6-[3-(((3-(trifluoromethyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.10 (s, 1H), 7.90 (m, 1H), 7.40 (m, 6H), 3.80 (d, 4H), 2.95 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{20}\text{N}_5\text{OF}_3$: 416.44 (MH^+).
- [0818] 3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.00 (s, 1H), 7.90 (d, 1H), 7.40 (m, 6H), 3.80 (m, 1H), 3.60 (m, 2H), 3.00 (s, 3H), 1.40 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{23}\text{N}_5\text{O}$: 362.49 (MH^+).
- [0819] 3-amino-6-[3-(((1R)-2-hydroxy-1-phenylethyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.00 (s, 1H), 7.90 (d, 1H), 7.40 (m, 7H), 3.80 (m, 2H), 3.60 (m, 3H), 3.00 (s, 3H); MS (EI) for $\text{C}_{21}\text{H}_{23}\text{N}_5\text{O}_2$: 378.48 (MH^+).
- [0820] methyl (2S)-[3-((3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)methyl)amino)(phenyl)ethanoate: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.00 (s, 1H), 7.90 (d, 1H), 7.40 (m, 7H), 3.80 (m, 6H), 3.00 (s, 3H); MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_5\text{O}_3$: 406.49 (MH^+).
- [0821] 3-amino-N-methyl-6-[3-([1-(phenylmethyl)pyrrolidin-3-yl]amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.60 (s, 1H), 8.10 (s, 1H), 7.90 (d, 1H), 7.38 (m, 9H), 3.80 (d, 2H), 3.60 (s, 2H), 3.40 (m, 1H), 2.90 (s, 3H), 2.82 (m, 1H), 2.60 (m, 2H), 2.40 (m, 1H), 2.18 (m, 1H), 1.70 (m, 1H); MS (EI) for $\text{C}_{24}\text{H}_{28}\text{N}_6\text{O}$: 417.47 (MH^+).
- [0822] 3-amino-6-(3-[[ethyl(phenylmethyl)amino]methyl]phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.60 (s, 1H), 7.90 (d, 1H), 7.82 (m, 1H), 7.30 (m, 8H), 3.61 (d, 2H), 3.59 (d, 2H), 2.90 (s, 3H), 2.50 (q, 2H), 1.10 (t, 3H); MS (EI) for $\text{C}_{22}\text{H}_{25}\text{N}_5\text{O}$: 376.44 (MH^+).
- [0823] 3-amino-N-methyl-6-[3-(((2S)-2-phenylcyclopropyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.60 (s, 1H), 7.99 (s, 1H),

7.85 (d, 1H), 7.40 (m, 2H), 7.15 (m, 5H), 6.90 (m, 2H), 3.90 (s, 2H), 2.90 (d, 3H), 2.38 (m, 1H), 1.85 (m, 1H), 1.10 (m, 1H), 1.00 (m, 1H); MS (EI) for $C_{22}H_{23}N_5O$: 374.43 (MH^+).

[0824] 3-amino-N-methyl-6-(3-([(1-methyl-1-phenylethyl)amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.61 (s, 1H), 7.99 (s, 1H), 7.85 (d, 1H), 7.60 (m, 2H), 7.40 (m, 7H), 3.42 (s, 2H), 3.00 (s, 3H), 1.60 (s, 6H); MS (EI) for $C_{22}H_{25}N_5O$: 376.42 (MH^+).

[0825] 3-amino-6-{3-[(9H-fluoren-9-ylamino)methyl]phenyl}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.61 (s, 1H), 7.99 (s, 1H), 7.85 (d, 1H), 7.70 (m, 5H), 7.40 (m, 7H), 7.20 (d, 1H), 3.41 (s, 2H), 3.00 (s, 3H); MS (EI) for $C_{26}H_{23}N_5O$: 422.39 (MH^+).

[0826] 3-amino-6-[3-([(1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.42 (m, 4H), 7.20 (m, 4H), 4.20 (m, 2H), 3.40 (s, 3H), 3.10 (m, 1H), 2.95 (m, 1H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390.25 (MH^+).

[0827] 3-amino-6-(3-([(5-bromo-2,3-dihydro-1H-inden-2-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.42 (m, 7H), 3.85 (s, 2H), 3.61 (m, 1H), 3.20 (m, 2H), 2.95 (s, 3H), 2.80 (m, 2H); MS (EI) for $C_{22}H_{22}N_5OBr$: 452.10 ($M+2$).

[0828] 3-amino-N-methyl-6-[3-([(1S,2R)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.42 (m, 4H), 7.20 (m, 4H), 4.20 (m, 2H), 3.40 (s, 3H), 3.40 (s, 3H), 3.10 (m, 1H), 3.00 (s, 3H), 2.95 (m, 1H); MS (EI) for $C_{23}H_{25}N_5O_2$: 404.27 (MH^+).

[0829] 3-amino-6-(3-([(5-fluoro-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.42 (m, 4H), 7.10 (m, 3H), 4.40 (s, 2H), 3.40 (m, 2H), 3.10 (m, 1H), 2.75 (m, 1H), 2.40 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OF$: 392.27 (MH^+).

[0830] 3-amino-6-[3-([(2S)-5-bromo-2,3-dihydro-1H-inden-2-yl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.15

(m, 2H), 7.35 (m, 6H), 3.80 (s, 2H), 3.58 (m, 1H), 3.16 (m, 2H), 2.82 (m, 3H), 2.70 (m, 2H); MS (EI) for $C_{22}H_{22}N_5OBr$: 454.14 (M+2).

- [0831] 3-amino-N-methyl-6-[3-({[(1R,2S)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.42 (m, 4H), 7.20 (m, 4H), 4.20 (m, 2H), 3.40 (s, 3H), 3.10 (m, 1H), 3.00 (s, 3H), 2.95 (m, 1H); MS (EI) for $C_{23}H_{25}N_5O_2$: 404.27 (MH⁺).
- [0832] 3-amino-N-methyl-6-(3-({[(3-oxo-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)pyrazine-2-carboxamide 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.15 (s, 1H), 7.90 (d, 1H), 7.42 (m, 4H), 7.20 (m, 4H), 4.50 (m, 1H), 3.90 (s, 2H), 3.40 (m, 1H), 3.02 (m, 1H), 3.00 (s, 3H), 2.60 (m, 1H); MS (EI) for $C_{22}H_{21}N_5O_2$: 388.19 (MH⁺).
- [0833] 3-amino-6-[3-({[(1S)-4-fluoro-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide 1H NMR (400 MHz, d4-MeOH): δ 8.64 (s, 1H), 8.10 (s, 1H), 7.85 (d, 1H), 7.40 (m, 3H), 7.20 (m, 3H), 6.90 (t, 1H), 4.40 (m, 1H), 3.90 (s, 2H), 3.10 (m, 1H), 2.98 (s, 3H), 2.80 (m, 1H), 2.42 (m, 1H), 2.10 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OF$: 392.16 (MH⁺).
- [0834] 3-amino-6-[3-({[(1S)-5,7-difluoro-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.64 (s, 1H), 8.10 (s, 1H), 7.85 (d, 1H), 7.40 (m, 3H), 7.20 (m, 2H), 4.50 (m, 1H), 3.90 (m, 2H), 3.20 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.42 (m, 1H), 2.15 (m, 1H); MS (EI) for $C_{22}H_{21}N_5OF_2$: 410.20 (MH⁺).
- [0835] 3-amino-N-methyl-6-[3-({[(1S)-5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl]amino}methyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.40 (m, 9H), 7.05 (m, 1H), 4.38 (m, 1H), 3.90 (m, 2H), 3.18 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.42 (m, 1H), 2.15 (m, 1H); MS (EI) for $C_{26}H_{25}N_5OS$: 456.14 (MH⁺).
- [0836] 3-amino-6-(3-({[(4-bromo-2,3-dihydro-1H-inden-1-yl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.40 (m, 6H), 7.16 (m, 1H), 4.42 (m, 1H), 3.90 (m, 2H), 3.15 (m, 1H), 3.00 (s, 3H), 2.82 (m, 1H), 2.42 (m, 1H), 2.02 (m, 1H); MS (EI) for $C_{22}H_{22}N_5OBr$: 454.01 (M+2).

- [0837] 3-amino-N-methyl-6-(3-([(4-phenyl-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.40 (m, 9H), 7.05 (m, 3H), 4.38 (m, 1H), 3.90 (m, 2H), 3.18 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.42 (m, 1H), 2.15 (m, 1H); MS (EI) for $\text{C}_{28}\text{H}_{27}\text{N}_5\text{O}$: 450.20 (MH^+).
- [0838] 3-amino-N-methyl-6-[3-([(4-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.42 (m, 7H), 7.27 (m, 2H), 7.11 (m, 1H), 4.38 (m, 1H), 3.90 (m, 2H), 3.30 (m, 1H), 3.00 (m, 1H), 2.95 (s, 3H), 2.42 (m, 1H), 2.12 (m, 1H); MS (EI) for $\text{C}_{26}\text{H}_{25}\text{N}_5\text{OS}$: 456.15 (MH^+).
- [0839] 3-amino-N-methyl-6-[3-([(2R)-6-phenyl-1,2,3,4-tetrahydronaphthalen-2-yl)amino]methyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.60 (m, 2H), 7.30 (m, 10H), 4.70 (m, 1H), 4.00 (m, 2H), 3.18 (m, 1H), 3.11 (m, 1H), 2.99 (s, 3H), 2.85 (m, 1H), 2.70 (m, 1H), 2.12 (m, 1H), 1.70 (m, 1H); MS (EI) for $\text{C}_{29}\text{H}_{29}\text{N}_5\text{O}$: 464.21 (MH^+).
- [0840] 3-amino-N-methyl-6-[3-([(5-[(2-morpholin-4-ylethyl)oxy]-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.40 (m, 5H), 6.80 (m, 2H), 4.35 (m, 1H), 4.10 (m, 2H), 3.90 (m, 2H), 3.70 (m, 4H), 3.10 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.83 (m, 2H), 2.60 (m, 4H), 2.40 (m, 1H), 2.05 (m, 1H); MS (EI) for $\text{C}_{28}\text{H}_{34}\text{N}_6\text{O}_3$: 503.17 (MH^+).
- [0841] 3-amino-6-[3-([(5-[(cyanomethyl)oxy]-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.40 (m, 5H), 6.90 (m, 2H), 4.95 (s, 2H), 4.35 (m, 1H), 3.90 (m, 2H), 3.10 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.40 (m, 1H), 2.05 (m, 1H); MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2$: 429.17 (MH^+).
- [0842] 3-amino-N-methyl-6-[3-([(5-(1H-1,2,4-triazol-1-yl)-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 9.10 (s, 1H), 8.70 (s, 1H), 8.17 (s, 2H), 7.90 (d, 1H), 7.60 (m, 4H), 7.40 (m, 3H), 4.40 (m, 1H), 3.95 (m, 2H), 3.16 (m, 1H), 3.00 (s, 3H), 2.92 (m, 1H), 2.50 (m, 1H), 2.03 (m, 1H); MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_8\text{O}$: 441.17 (MH^+).
- [0843] 3-amino-6-(3-([(4-hydroxy-2,3-dihydro-1H-inden-1-yl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d4-MeOH): δ 8.70 (s, 1H), 8.17 (s,

1H), 7.92 (d, 1H), 7.40 (m, 2H), 7.20 (m, 2H), 6.65 (m, 3H), 4.40 (br, OH, 1H), 4.21 (m, 1H), 3.90 (m, 2H), 3.10 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.40 (m, 1H), 2.05 (m, 1H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390.16 (MH^+).

[0844] 3-amino-N-methyl-6-{3-[[4-[(2-morpholin-4-ylethyl)oxy]-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.40 (m, 4H), 7.20 (m, 1H), 7.00 (m, 1H), 6.80 (m, 1H), 4.35 (m, 1H), 4.10 (m, 2H), 3.90 (m, 2H), 3.70 (m, 4H), 3.10 (m, 1H), 3.00 (s, 3H), 2.85 (m, 1H), 2.83 (m, 2H), 2.60 (m, 4H), 2.40 (m, 1H), 2.05 (m, 1H); MS (EI) for $C_{28}H_{34}N_6O_3$: 503.20 (MH^+).

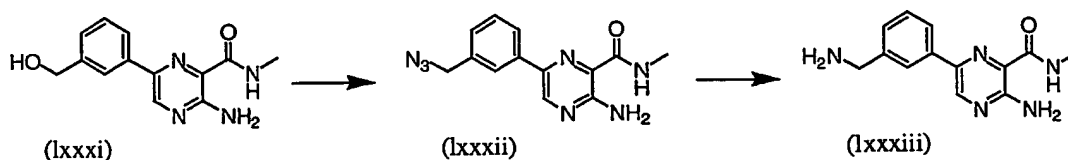
[0845] 3-amino-6-(3-{[(4-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.17 (s, 1H), 7.92 (d, 1H), 7.42 (m, 4H), 7.10 (m, 1H), 6.95 (m, 1H), 6.61 (m, 1H), 4.31 (m, 1H), 3.90 (m, 2H), 3.10 (m, 1H), 3.00 (s, 3H), 2.76 (m, 1H), 2.40 (m, 1H), 2.05 (m, 1H); MS (EI) for $C_{22}H_{23}N_5O_2$: 390.16 (MH^+).

[0846] 3-amino-6-[3-(1H-imidazol-1-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 8.81 (s, 1H), 8.76 (m, 1H), 8.10 (m, 2H), 7.80 (s, 1H), 7.66 (br s, 2H), 7.41 (t, 1H), 7.24 (s, 1H), 7.17 (d, 1H), 6.89 (s, 1H), 5.24 (s, 2H), 2.86 (d, 3H); MS (EI) for $C_{16}H_{16}N_6O$: 309.2 (MH^+).

Example 53

[0847] Scheme 18 shows how alcohols were converted to amines to make compounds according to formula I. In this example, alcohol (lxxxi) is converted to the corresponding azide (lxxxii), which was reduced to the corresponding amine (lxxxiii). Amines (lxxxiii) were, for example, acylated, subjected to reductive amination, and the like to make compounds according to formula I.

Scheme 18



3-Amino-6-[3-(aminomethyl)phenyl]-N-methylpyrazine-2-carboxamide

- [0848] 3-Amino-6-(3-hydroxymethyl-phenyl)-pyrazine-2-carboxylic acid methylamide (1.00 g, 3.87 mmol) was dissolved in THF (20 mL) followed by the addition of diphenylphosphoryl azide (0.63 mL, 5.42 mmol) and DBU (0.43 mL, 5.42 mmol). It was stirred at room temperature overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography (50% Ethylacetate/hexanes) to give 3-amino-6-(3-azidomethyl-phenyl)-pyrazine-2-carboxylic acid methylamide, 728 mg (66.4 % yield): MS (EI) for C₁₃H₁₃N₇O₀: 284.12 (MH⁺).
- [0849] 3-Amino-6-(3-azidomethyl-phenyl)-pyrazine-2-carboxylic acid methylamide was dissolved in the mixture of THF (15 mL) and H₂O (1.5 mL) followed by the addition of triphenylphosphine (1.07 g, 4.03 mmol) and it was stirred at room temperature overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate and filtered, concentrated and purified by column chromatography (50% Ethylacetate/hexanes) to give 3-amino-6-(3-aminomethyl-phenyl)-pyrazine-2-carboxylic acid methylamide, 564 mg (85.0% yield): MS (EI) for C₁₃H₁₅N₅O: 258.13 (MH⁺).
- [0850] 3-Amino-N-methyl-6-(3-([(phenylcarbonyl)amino]methyl)phenyl)pyrazine-2-carboxamide 3-Amino-6-(3-aminomethyl-phenyl)-pyrazine-2-carboxylic acid methylamide (10.0 mg, 0.04 mmol) was dissolved in acetonitrile (2 mL) followed by the addition of TEA (11.2 μL, 0.08 mmol) and benzoyl chloride (3.50 μL, 0.04 mmol). It was stirred at room temperature overnight. The reaction mixture was poured into water (10 mL), and extracted with ethyl acetate (2 × 10 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography (50% Ethylacetate/hexanes) to give 3-amino-6-[3-(benzoylamino-methyl)-phenyl]-pyrazine-2-carboxylic acid methylamide, 8.70 mg (85.0% yield): ¹H NMR (400 MHz, CDCl₃): δ 8.60 (s, 1H), 8.00 (br, 1H), 7.80 (m, 4H), 7.52 (m, 4H), 7.20 (s, 1H), 6.50 (br, 1H), 4.80 (m, 2H), 3.00 (d, 3H); MS (EI) for C₂₀H₁₉N₅O₂: 362.41 (MH⁺).
- [0851] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.
- [0852] 3-Amino-6-[3-([(4-chlorophenyl)carbonyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, CDCl₃): δ 8.60 (s, 1H), 8.00 (br,

1H), 7.80 (m, 3H), 7.52 (m, 4H), 7.20 (s, 1H), 6.42 (br, 1H), 4.78 (m, 2H), 3.00 (d, 3H); MS (EI) for $C_{20}H_{18}ClN_5O_2$: 396.36 (MH^+).

[0853] 3-Amino-6-[3-(((2,5-difluorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.10 (br, 1H), 8.80 (s, 1H), 8.20 (m, 2H), 7.40 (m, 5H), 4.60 (m, 2H), 2.80 (d, 3H); MS (EI) for $C_{20}H_{17}F_2N_5O_2$: 398.43 (MH^+).

[0854] 3-amino-N-methyl-6-(3-([(naphthalen-2-ylcarbonyl)amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.30 (br, 1H), 8.82 (s, 1H), 8.80 (br, 1H), 8.50 (s, 1H), 8.00 (m, 5H), 7.60 (m, 2H), 7.40 (m, 2H), 4.60 (m, 2H), 2.80 (d, 3H); MS (EI) for $C_{24}H_{21}N_5O_2$: 412.27 (MH^+).

[0855] 3-amino-6-[3-([(3,4-dichlorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.40 (br, 1H), 8.82 (s, 1H), 8.80 (br, 1H), 8.20 (s, 1H), 8.10 (m, 2H), 7.90 (d, 1H), 7.80 (d, 1H), 7.40 (m, 1H), 7.38 (m, 1H), 4.60 (m, 2H), 2.80 (d, 3H); MS (EI) for $C_{20}H_{17}Cl_2N_5O_2$: 430.36 (MH^+).

[0856] 3-amino-N-methyl-6-(3-([(phenylacetyl)amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.71 (t, 1H), 8.01 (m, 3H), 7.40 (m, 1H), 7.22 (m, 1H), 7.18 (m, 3H), 6.90 (m, 2H), 4.60 (d, 2H), 4.40 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{21}H_{21}N_5O_2$: 376.46 (MH^+).

[0857] 3-amino-6-[3-([(4-chlorophenyl)acetyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.71 (t, 1H), 8.01 (m, 2H), 7.40 (m, 1H), 7.22 (m, 1H), 7.18 (m, 3H), 6.90 (m, 2H), 4.60 (d, 2H), 4.40 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{21}H_{20}ClN_5O_2$: 410.45 (MH^+).

[0858] 3-amino-6-(3-([(biphenyl-4-ylcarbonyl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, $CDCl_3$): δ 8.60 (s, 1H), 7.98 (br, 1H), 7.85 (m, 3H), 7.70 (m, 1H), 7.60 (m, 4H), 7.40 (m, 3H), 7.22 (s, 2H), 6.50 (br, 1H), 4.80 (d, 2H), 3.00 (d, 3H); MS (EI) for $C_{26}H_{23}N_5O_2$: 438.51 (MH^+).

[0859] 3-amino-6-[3-([(2-chloro-5-(trifluoromethyl)phenyl)carbonyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, $CDCl_3$): δ 8.60 (s, 1H), 7.80 (m, 3H), 7.40 (m, 3H), 7.22 (s, 2H), 6.60 (br, 1H), 4.80 (d, 2H), 3.00 (d, 3H); MS (EI) for $C_{21}H_{17}ClF_3N_5O_2$: 464.42 (MH^+).

- [0860] N-[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)methyl]-1H-indole-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 11.60 (s, 1H), 8.80 (s, 1H), 8.70 (m, 1H), 8.40 (m, 1H), 8.00 (m, 4H), 7.80 (m, 3H), 7.40 (m, 3H), 7.10 (m, 2H), 4.60 (d, 2H), 2.80 (d, 3H); MS (EI) for $\text{C}_{22}\text{H}_{20}\text{N}_6\text{O}_2$: 401.48 (MH^+).
- [0861] 3-amino-6-[3-({[(3-hydroxypyridin-2-yl)carbonyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 12.00 (s, 1H), 8.60 (s, 1H), 8.40 (br, 1H), 7.80 (m, 3H), 7.28 (m, 5H), 7.10 (m, 2H), 4.75 (d, 2H), 3.00 (d, 3H), 1.60 (br, 2H); MS (EI) for $\text{C}_{19}\text{H}_{18}\text{N}_6\text{O}_3$: 379.46 (MH^+).
- [0862] 3-amino-6-(3-({[(1H-indole-3-ylacetyl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, CDCl_3): δ 10.85 (s, 1H), 8.80 (m, 1H), 8.40 (m, 1H), 8.00 (m, 2H), 7.50 (d, 2H), 7.40 (m, 3H), 7.00 (m, 2H), 4.40 (d, 2H), 3.60 (s, 3H), 3.40 (s, 2H); MS (EI) for $\text{C}_{23}\text{H}_{22}\text{N}_6\text{O}_2$: 414.18 (MH^+).
- [0863] 3-amino-N-methyl-6-[3-({[(4-(1-methylethyl)phenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 9.12 (br, 1H), 8.90 (s, 1H), 8.80 (br, 1H), 8.50 (s, 1H), 8.38 (d, 1H), 7.80 (d, 1H), 7.50 (t, 1H), 7.20 (m, 4H), 4.50 (d, 2H), 2.80 (d, 3H), 1.20 (m, 6H), 0.80 (m, 1H); MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_5\text{O}_2$: 404.52 (MH^+).
- [0864] 3-amino-N-methyl-6-(3-({[(4-(1-methylethyl)phenyl)oxy]acetyl]amino}methyl)phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 8.90 (m, 2H), 8.72 (t, 1H), 8.05 (m, 2H), 7.40 (m, 1H), 7.20 (m, 3H), 6.90 (d, 2H), 4.60 (s, 2H), 4.40 (d, 2H), 2.80 (d, 3H), 1.20 (m, 6H), 0.80 (m, 1H); MS (EI) for $\text{C}_{24}\text{H}_{27}\text{N}_5\text{O}_3$: 434.51 (MH^+).
- [0865] 3-amino-6-[3-({[(4-aminophenyl)thio]acetyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 8.90 (m, 2H), 8.70 (t, 1H), 8.00 (m, 2H), 7.40 (m, 1H), 7.18 (m, 3H), 6.50 (d, 2H), 5.20 (s, 2H), 4.40 (d, 2H), 2.80 (d, 3H); MS (EI) for $\text{C}_{21}\text{H}_{22}\text{N}_6\text{O}_2\text{S}$: 423.44 (MH^+).
- [0866] 3-amino-6-[3-({[(4-iodophenyl)carbonyl]amino}methyl)phenyl]-N-methylpyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 9.20 (m, 1H), 8.90 (m, 2H), 8.12 (m, 2H), 7.80 (m, 2H), 7.70 (m, 2H), 7.40 (m, 2H), 4.50 (d, 2H), 2.80 (d, 3H); MS (EI) for $\text{C}_{20}\text{H}_{18}\text{N}_5\text{O}_2\text{I}$: 488.35 (MH^+).
- [0867] 3-amino-N-methyl-6-[3-({[(4-pentylphenyl)carbonyl]amino}methyl)phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 9.00 (m, 1H), 8.90 (m, 2H),

8.12 (m, 2H), 7.80 (m, 2H), 7.40 (m, 4H), 4.60 (d, 2H), 3.40 (d, 3H), 1.60 (m, 2H), 1.20 (m, 5H), 0.80 (m, 4H); MS (EI) for $C_{25}H_{29}N_5O_2$: 432.55 (MH^+).

[0868] 3-amino-6-(3-([(1,3-benzodioxol-5-ylcarbonyl)amino]methyl)phenyl)-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.78 (m, 2H), 8.50 (br, 1H), 8.08 (d, 1H), 7.98 (s, 1H), 7.42 (t, 1H), 7.20 (d, 1H), 6.80 (m, 3H), 4.40 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{21}H_{19}N_5O_4$: 406.15 (MH^+).

[0869] 3-amino-6-{3-([(4-(1,1-dimethylethyl)phenyl)methyl]amino)carbonyl]phenyl}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.10 (br, 1H), 8.90 (s, 1H), 8.80 (br, 1H), 8.50 (s, 1H), 8.40 (d, 1H), 7.82 (d, 1H), 7.60 (t, 1H), 7.35 (m, 4H), 4.50 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{24}H_{27}N_5O_2$: 418.52 (MH^+).

[0870] 3-amino-N-methyl-6-(3-([(1-[5-(trifluoromethyl)pyrimidin-2-yl]piperidin-4-yl)carbonyl]amino)methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.80 (m, 2H), 8.62 (m, 1H), 8.40 (t, 1H), 8.00 (m, 2H), 7.40 (t, 1H), 7.20 (d, 1H), 7.00 (d, 1H), 4.60 (d, 2H), 4.38 (d, 2H), 2.98 (t, 2H), 2.80 (d, 3H), 1.80 (m, 2H), 1.60 (m, 2H), 1.20 (m, 1H); MS (EI) for $C_{24}H_{25}N_8O_2F_3$: 515.51 (MH^+).

[0871] 3-amino-N-methyl-6-{3-([(4-(1-methylethyl)phenyl)carbonyl]amino)methyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.00 (br, 1H), 8.90 (m, 2H), 8.12 (m, 2H), 7.82 (d, 1H), 7.35 (m, 4H), 4.60 (d, 2H), 2.90 (m, 1H), 2.80 (d, 3H), 1.20 (d, 6H); MS (EI) for $C_{23}H_{25}N_5O_2$: 404.50 (MH^+).

[0872] 3-amino-6-[3-([(4-butylphenyl)methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.10 (br, 1H), 8.90 (m, 2H), 8.50 (s, 1H), 8.40 (d, 1H), 7.82 (d, 1H), 7.60 (m, 1H), 7.20 (m, 4H), 4.60 (d, 2H), 2.80 (d, 3H), 2.60 (m, 2H), 1.60 (m, 2H), 1.32 (m, 2H), 0.90 (m, 3H); MS (EI) for $C_{24}H_{27}N_5O_2$: 418.49 (MH^+).

[0873] 3-amino-N-methyl-6-{3-([(4-(1,2,3-thiadiazol-4-yl)phenyl)methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.60 (s, 1H), 9.20 (br, 1H), 8.90 (m, 2H), 8.60 (s, 1H), 8.40 (d, 1H), 8.18 (d, 2H), 7.98 (d, 1H), 7.58 (m, 3H), 4.60 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{22}H_{19}N_7O_2S$: 446.42 (MH^+).

[0874] 3-amino-N-methyl-6-{3-([(4-(2-thienyl)phenyl)methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (s, 1H), 8.90

(m, 2H), 8.60 (s, 1H), 8.40 (d, 1H), 7.98 (d, 1H), 7.58 (m, 7H), 7.18 (d, 1H), 4.60 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{24}H_{21}N_5O_2S$: 444.46 (MH^+).

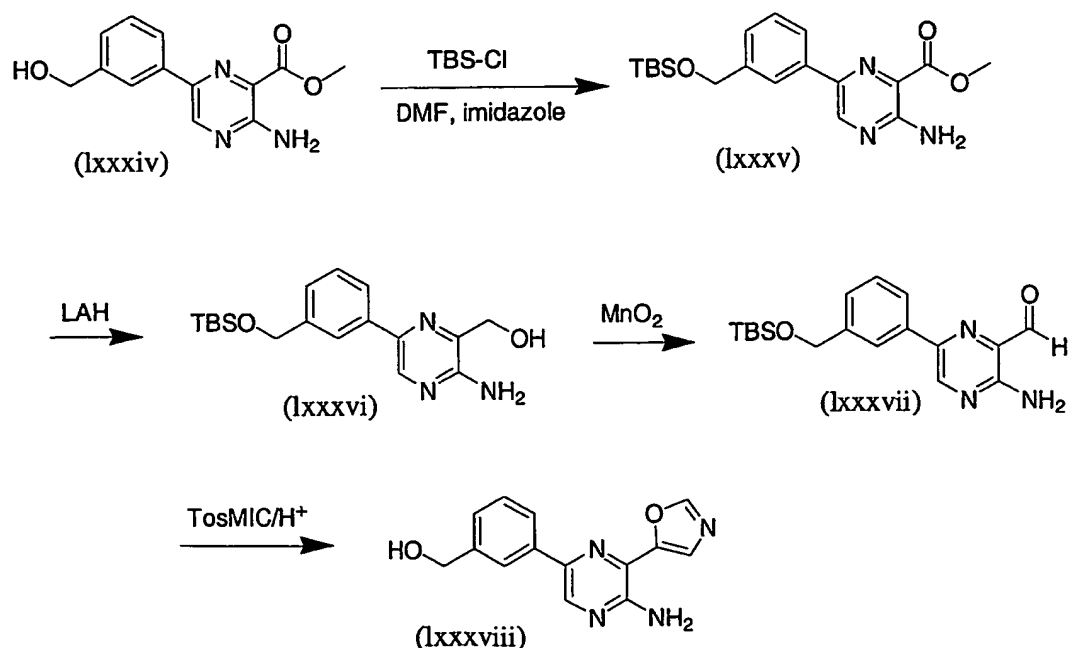
[0875] 3-amino-6-{3-[[[4-(1,1-dimethylethyl)phenyl]carbonyl]amino)methyl]phenyl}-N-methylpyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.10 (s, 1H), 8.90 (m, 2H), 8.00 (m, 4H), 7.40 (m, 4H), 4.60 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{24}H_{27}N_5O_2$: 418.51 (MH^+).

[0876] 3-amino-N-methyl-6-{3-[[[4-(1H-pyrazol-1-yl)phenyl]methyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (br, 1H), 8.90 (m, 2H), 8.50 (m, 3H), 7.60 (m, 7H), 6.50 (s, 1H), 4.60 (d, 2H), 2.80 (d, 3H); MS (EI) for $C_{23}H_{21}N_7O_2$: 428.48 (MH^+).

Example 54

[0877] Scheme 19 depicts a strategy for making exemplary compounds according to formula I where A is an oxazole. Compound (lxxxiv) is protected as the TBS-ether (lxxxv). The ester function of (lxxxv) is reduced to the corresponding alcohol, (lxxxvi), and reoxidized to the corresponding aldehyde, (lxxxvii). Aldehyde (lxxxvii) was converted to oxazole (lxxxviii) as described below.

Scheme 19



[0878] {3-[5-amino-6-(1,3-oxazol-5-yl)pyrazin-2-yl]phenyl}methanol: Methyl 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carboxylate (200mg, 0.77 mmol) was dissolved in DMF followed by addition of *tert*-butylsilyl chloride (0.14g, 0.93 mmol) and imidazole (63 mg, 9.25 mmol). This mixture was stirred at room temperature under nitrogen for overnight. The reaction mixture was poured into water (75 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate and filtration, concentration to give crude silyl ether product, 170 mg (59.0%). It was dissolve in THF followed by addition of lithium aluminum hydride (1.0 M solution in THF, 0.9 ml, about 2.0 equiv.) and it was stirred at room temperature for 5 hrs. It was directly subjected to oxidation reaction with MnO₂. Crude compounds was dissolved in acetone followed by the addition of MnO₂ (730 mg, 20 equiv.). It was refluxed for 1hr then dried over anhydrous sodium sulfate and filtration, concentration to give crude aldehyde, 94.0 mg (66.0%). The aldehyde was dissolved in MeOH (20 ml), followed by tosylmethylisocyanide (56.0 mg, 2.87 mmol) and K₂CO₃ (40.0 mg, 2.87 mmol). This reaction mixture was refluxed for 4 hrs. and then followed usual aqueous work up and it was treated with HCl to give the desired oxazole, 62.0 mg (84.4%). ¹H NMR (400 MHz, d₄-MeOH): δ 8.70 (s, 1H), 8.01 (s, 1H), 7.80 (s, 1H), 7.20 (m, 4H), 4.70 (m, 2H); MS (EI) for C₁₄H₁₂N₄O₂: 267.14 (MH⁺).

Example 55

5-(3-([(1S)-2,3-dihydro-1H-inden-1-yl]amino)methyl)phenyl)-3-(1,3-oxazol-5-yl)pyrazin-2-amine

[0879] Using the {3-[5-Amino-6-(1,3-oxazol-5-yl)pyrazin-2-yl]phenyl}methanol, Mitsunobu reactions were carried out as outlined in Scheme 15 above to make compounds according to formula I.

[0880] To a solution of oxazole (Ixxxviii) (I (62.0 mg, 2.31 mmol) in benzene (0.5 M solution) was added N-[(1S)-2,3-dihydro-1H-inden-1-yl]2,4-dinitrobenzenesulfonamide (42.0 mg, 1.15 mmol), triphenylphosphine (57.7 mg, 2.19 mmol) and DEAD (0.034 mL, 2.31 mmol) and the mixture was allowed to stir at room temperature overnight. The solvent was removed and directly subjected to column chromatography purification (50% ethylacetate/hexanes) to give the intermediate sulfonamide, 123 mg (84.8 % yield), which was treated with *n*-propylamine (20 equiv.) and purified by MPLC to give 5-(3-([(1S)-

2,3-dihydro-1H-inden-1-ylamino)methyl}phenyl)-3-(1,3-oxazol-5-yl)pyrazin-2-amine, 100 mg (81.3% yield): ^1H NMR (400 MHz, d₄-MeOH): δ 8.58 (s, 1H), 8.40 (s, 1H), 8.00 (s, 1H), 7.82 (d, 1H), 7.80 (s, 1H), 7.40 (m, 4H), 7.20 (4H), 4.35 (m, 1H), 3.90 (m, 2H), 3.10 (m, 1H), 2.80 (m, 1H), 2.40 (m, 1H), 1.98 (m, 1H); MS (EI) for C₂₃H₂₁N₅O: 384.15 (MH⁺).

[0881] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.

[0882] 1-[3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino)methyl}phenyl]pyrazin-2-yl)ethanone: ^1H NMR (400 MHz, d₄-MeOH): δ 8.70 (s, 1H), 8.40 (s, 1H), 7.98 (d, 1H), 7.80 (m, 2H), 7.40 (m, 3H), 7.20 (m, 3H), 4.21 (m, 1H), 3.82 (m, 2H), 3.10 (m, 1H), 2.81 (m, 1H), 2.61 (m, 3H), 2.38 (m, 1H), 1.97 (m, 1H); MS (EI) for C₂₂H₂₂N₄O: 359.15 (MH⁺).

[0883] [3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino)methyl}phenyl]pyrazin-2-yl)(phenyl)methanone: ^1H NMR (400 MHz, d₄-MeOH): δ 8.80 (s, 1H), 8.00 (d, 2H), 7.98 (s, 1H), 7.80 (m, 1H), 7.40 (m, 8H), 7.20 (m, 4H), 4.40 (m, 1H), 3.90 (m, 2H), 3.10 (m, 1H), 2.85 (m, 1H), 2.40 (m, 1H), 2.05 (m, 1H); MS (EI) for C₂₇H₂₄N₄O: 421.20 (MH⁺).

[0884] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino)methyl}phenyl)-3-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine: ^1H NMR (400 MHz, d₄-MeOH): δ 10.1 (br, 1H), 9.00 (m, 3H), 8.70 (d, 1H), 8.30 (d, 1H), 7.91 (d, 1H), 7.40 (m, 7H), 4.81 (br, 1H), 4.35 (br, 1H), 4.71 (m, 1H), 3.21 (m, 1H), 2.91 (m, 1H), 2.51 (m, 1H); MS (EI) for C₂₇H₂₄N₈: 461.16 (MH⁺).

[0885] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino)methyl}phenyl)-3-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine: ^1H NMR (400 MHz, d₄-MeOH): δ 8.70 (s, 1H), 8.40 (br, 1H), 8.21 (m, 1H), 7.60 (m, 3H), 7.40 (m, 3H), 4.35 (s, 2H), 3.71 (m, 6H), 3.12 (m, 2H), 2.70 (m, 1H), 2.40 (m, 2H), 2.18 (m, 3H); MS (EI) for C₂₇H₃₀N₈: 467.41 (MH⁺).

[0886] 5-{3-[(2,3-dihydro-1H-inden-1-ylamino)methyl]phenyl}-3-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine: ^1H NMR (400 MHz, d₄-MeOH): δ 8.60 (d, 2H), 8.18 (d, 1H), 7.79 (d, 1H), 7.60 (m, 2H), 7.40 (m, 3H), 4.40 (s, 2H), 3.60 (m, 6H), 3.12 (m, 1H), 3.05 (m, 1H), 2.67 (m, 1H), 2.42 (m, 1H), 2.18 (m, 1H), 2.10 (m, 3H); MS (EI) for C₂₇H₃₀N₈: 467.18 (MH⁺).

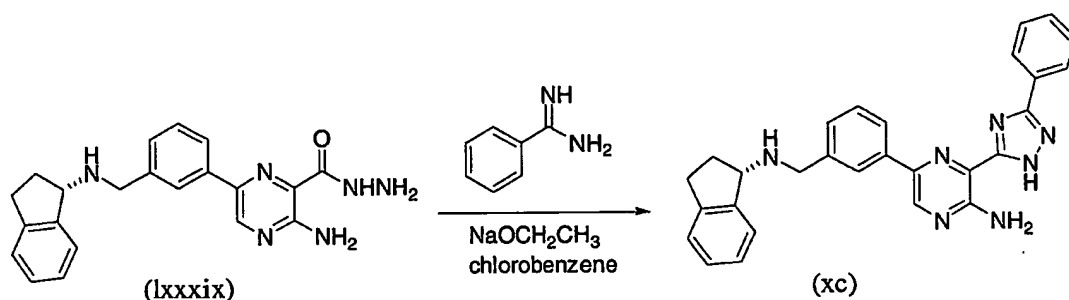
[0887] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.60 (d, 2H), 8.18 (d, 1H), 7.79 (d, 1H), 7.60 (m, 2H), 7.40 (m, 3H), 4.40 (s, 2H), 3.60 (m, 5H), 3.30 (m, 1H), 3.18 (m, 1H), 3.02 (m, 1H), 2.62 (m, 1H), 2.41 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $\text{C}_{27}\text{H}_{30}\text{N}_8$: 467.33(MH^+).

[0888] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-{3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.60 (d, 2H), 8.18 (d, 1H), 7.79 (d, 1H), 7.60 (m, 2H), 7.40 (m, 3H), 4.40 (s, 2H), 3.60 (m, 5H), 3.30 (m, 1H), 3.18 (m, 1H), 3.02 (m, 1H), 2.62 (m, 1H), 2.41 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $\text{C}_{27}\text{H}_{30}\text{N}_8$: 467.33 (MH^+).

Example 56

[0889] Scheme 20 depicts a strategy for making exemplary compounds according to formula I where A is a triazole. Compound (lxxxix) is converted to the corresponding triazole, (xc), for example, as described below.

Scheme 20



[0890] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-(3-phenyl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine: A solution of sodium ethoxide (19 mg, 0.27 mmol) in anhydrous ethanol (2 mL) was added to a solution of benzenecarboximidamide hydrochloride hydrate (31.35 mg, 0.2 mmol) in anhydrous ethanol (3 mL) at room temperature. The mixture was stirred at room temperature for 45 min. and filtered. To the ethanol filtrate was added 3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)pyrazine-2-carbohydrazide (50mg, 0.13 mmol) and the resulting solution was heated to reflux overnight. The solvent was removed and purified by MPLC to give compound 1 (12.5 mg 21.3% yield): ^1H NMR (400 MHz, $\text{d}_6\text{-DMSO}$): δ 10.0 (s,

1H), 9.00 (m, 3H), 8.70 (d, 1H), 8.30 (d, 1H), 7.70 (d, 1H), 7.35 (m, 8H), 4.81 (br, 1H), 4.35 (br, 2H), 3.62 (m, 1H), 3.55 (m, 1H), 3.20 (m, 1H), 2.95 (m, 1H); MS (EI) for $C_{28}H_{25}N_7$: 460.10 (MH^+).

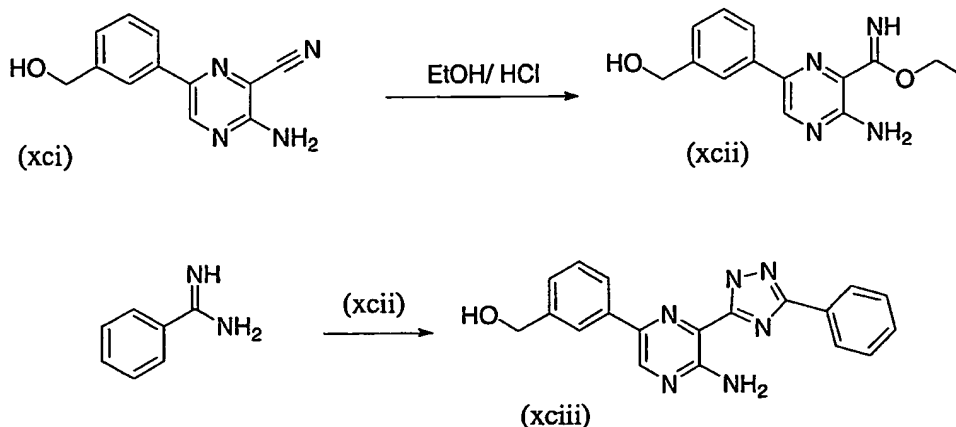
[0891] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.

[0892] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine: 1H NMR (400 MHz, d_6 -DMSO): δ 10.0 (br, 1H), 9.00 (m, 3H), 8.70 (d, 1H), 8.30 (d, 1H), 7.90 (d, 1H), 7.40 (m, 7H), 4.81 (br, 1H), 4.40 (br, 2H), 3.70 (m, 1H), 3.55 (m, 1H), 3.25 (m, 1H), 2.95 (m, 1H), 2.50 (m, 1H); MS (EI) for $C_{27}H_{24}N_8$: 461.12 (MH^+).

Example 57

[0893] Scheme 21 depicts a strategy for making imidoates according to formula I. Nitrile (xci) is converted to the corresponding imidoate, (xcii), for example, as described below. Such imidoates are generally used as intermediates to make, for example, compounds according to formula I where A is a triazole, or oxadiazole, and the like as one skilled in the art could readily perform. For example addition of benzenecarboximidamide to imidoate (xcii) under appropriate conditions (e.g. heat and dehydrative conditions) would give the corresponding triazole derivative (xciii). Some exemplary compounds are listed in this example.

Scheme 21



- [0894] Ethyl 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carboximidoate: A suspension of 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carbonitrile (500 mg, 2.21 mmol) in ethanol was cooled by ice bath. Dry hydrogen chloride gas was bubbled into the reaction mixture for 2 or 3 hours. After gas addition was stopped, the solution was placed in a refrigerator overnight. The solution was filtered and evaporated to afforded compound 1 (452 mg, 75.3% yield): ^1H NMR (400 MHz, d_6 -DMSO): δ 9.00 (s, 1H), 8.15 (s, 1H), 8.00 (m, 1H), 7.42 (m, 3H), 4.80 (q, 2H), 4.62 (s, 2H), 1.60 (t, 1H); MS (EI) for $\text{C}_{14}\text{H}_{16}\text{N}_4\text{O}$: 273.23 (MH^+).
- [0895] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.
- [0896] 2-({3-Amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl}carbonyl)-N-phenylhydrazine carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.42 (s, 1H), 8.20 (m, 3H), 7.40 (m, 8H), 6.95 (m, 1H), 5.20 (t, 1H), 4.60 (m, 2H); MS (EI) for $\text{C}_{19}\text{H}_{18}\text{N}_6\text{O}_3$: 379.13 (MH^+).
- [0897] 3-Amino-N-hydroxy-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carboximidamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.23 (s, 1H), 8.60 (s, 1H), 8.00 (m, 2H), 7.40 (m, 2H), 6.15 (s, 1H), 5.21 (t, 1H), 4.60 (m, 2H); MS (EI) for $\text{C}_{12}\text{H}_{13}\text{N}_5\text{O}_2$: 260.21 (MH^+).
- [0898] {3-[5-Amino-6-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.60 (s, 1H), 8.92 (m, 3H), 8.20 (m, 2H), 7.90 (m, 2H), 7.40 (m, 2H), 5.21 (t, 1H), 4.60 (m, 2H); MS (EI) for $\text{C}_{18}\text{H}_{15}\text{N}_7\text{O}$: 346.11 (MH^+).
- [0899] (3-{5-Amino-6-[3-(4-chlorophenyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.70 (s, 1H), 8.20 (m, 3H), 7.80 (m, 3H), 7.40 (m, 3H), 4.60 (m, 2H), 4.40 (s, 1H); MS (EI) for $\text{C}_{19}\text{H}_{15}\text{N}_6\text{OCl}$: 379.07 (MH^+).
- [0900] {3-[5-amino-6-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.80 (m, 5H), 8.10 (m, 3H), 7.50 (m, 2H), 7.40 (m, 3H), 4.72 (s, 2H) MS (EI) for $\text{C}_{18}\text{H}_{15}\text{N}_7\text{O}$: 346.24 (MH^+).
- [0901] {3-[5-amino-6-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.50 (s, 5H), 8.05 (s, 3H), 7.90

(m, 1H), 7.40 (m, 2H), 4.60 (s, 2H), 2.30 (m, 4H), 1.21 (m, 5H); MS (EI) for $C_{18}H_{12}N_7O$: 352.28 (MH^+).

[0902] {3-[5-amino-6-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: 1H NMR (400 MHz, d_6 -DMSO): δ 9.50 (dd, 2H), 8.78 (s, 1H), 8.20 (m, 2H), 7.40 (m, 2H), 4.60 (s, 2H), 3.32 (m, 4H), 1.67 (m, 5H); MS (EI) for $C_{18}H_{12}N_7O$: 352.28 (MH^+).

[0903] {3-[5-amino-6-(3-pyrrolidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: 1H NMR (400 MHz, d_6 -DMSO): δ 8.60 (s, 1H), 8.18 (s, 1H), 8.00 (d, 1H), 7.45 (m, 2H), 4.70 (s, 2H), 3.97 (m, 1H), 3.78 (m, 2H), 3.50 (m, 2H), 2.60 (m, 1H), 2.41 (m, 1H); MS (EI) for $C_{17}H_{19}N_7O$: 338.30 (MH^+).

[0904] {3-[5-amino-6-(3-pyridin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: 1H NMR (400 MHz, d_6 -DMSO): δ 8.91 (m, 2H), 8.56 (d, 1H), 8.40 (t, 1H), 8.18 (m, 2H), 7.81 (t, 1H), 7.42 (m, 2H), 4.70 (s, 2H); MS (EI) for $C_{18}H_{15}N_7O$: 346.22 (MH^+).

[0905] {3-[5-amino-6-(3-piperidin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: 1H NMR (400 MHz, d_6 -DMSO): δ 9.85 (m, 1H), 9.40 (m, 1H), 8.80 (s, 1H), 8.18 (m, 2H), 7.42 (m, 2H), 4.62 (s, 2H), 3.40 (m, 3H), 1.62 (m, 6H); MS (EI) for $C_{18}H_{21}N_7O$: 461.16 (MH^+).

[0906] {3-[5-amino-6-(3-morpholin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol: 1H NMR (400 MHz, d_6 -DMSO): δ 9.85 (m, 1H), 9.60 (m, 1H), 8.80 (s, 1H), 8.18 (m, 2H), 7.42 (m, 2H), 5.12 (d, 1H), 4.62 (s, 2H); MS (EI) for $C_{17}H_{19}N_7O_2$: 354.15 (MH^+).

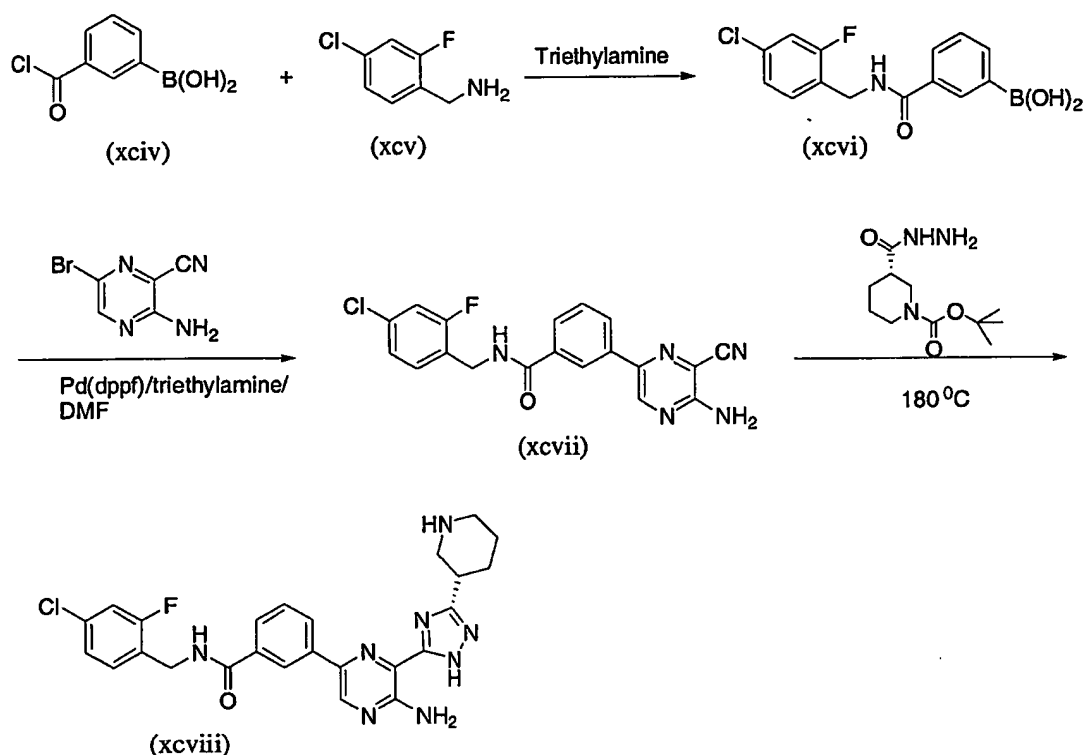
[0907] [3-(5-amino-6-{3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methanol: 1H NMR (400 MHz, d_4 -MeOH): δ 9.60 (s, 1H), 8.20 (s, 1H), 8.10 (m, 2H), 4.72 (s, 2H), 3.62 (m, 2H), 3.54 (m, 2H), 3.10 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $C_{18}H_{21}N_7O$: 352.17 (MH^+).

[0908] [3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methanol: 1H NMR (400 MHz, d_4 -MeOH): δ 9.60 (s, 1H), 8.20 (s, 1H), 8.10 (m, 2H), 4.72 (s, 2H), 3.62 (m, 2H), 3.54 (m, 2H), 3.10 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $C_{18}H_{21}N_7O$: 352.17 (MH^+).

Example 58

[0909] Scheme 22 depicts another strategy for making exemplary compounds according to formula I where A is a triazole. Boronic acid-acyl choride intermediate (xciv) is coupled with amine (xcv) to give amide (xcvi). Intermediate (xcvi) is coupled to a bromo-pyrazine, for example, to give intermediate (xcvii). The nitrile group of (xcvii) is converted to the corresponding triazole to yield (xcviii), for example, as more fully described below. Intermediate (xcvii) can also be used to make other heterocycles for A, according to formula I. Included in this example are other various triazole syntheses that generally start from a corresponding nitrile precursor.

Scheme 22



3-(5-amin-6-cyano-pyrazin-2-yl)-N-[(4-chloro-2-fluorophenyl)methyl]benzamide

[0910] 4-Carbonyl chloride phenylboronic acid (350 mg, 1.90 mmol) was added to THF followed by addition of 4-chloro-2-fluorobenzylamine (466 mg, 2.37 mmol) and triethylamine (400 μ l, 2.85 mmol). This mixture was stirred at room temperature under nitrogen overnight. The reaction mixture was poured into water (75 mL), and extracted with ethyl acetate (2 \times 50 mL) then dried over anhydrous sodium sulfate and filtration,

concentration to give crude reduction product, 525 mg (90.0% yield). Subsequently it was dissolved in DMF followed by addition of 3-amino-6-bromo-pyrazine-2-carbonitrile (169 mg, 0.85 mmol), Pd(dppf)CH₂Cl₂ (69.2 mg, 10 mol%) and triethylamine (118 μ L, 2.12 mmol). It was heated to 80 °C for 16 hrs and quenched with H₂O and extracted with ethylacetate then dried over anhydrous sodium sulfate and filtration, concentration to give crude reduction product (275 mg, 85.0% yield): MS (EI) for C₁₉H₁₃ClFN₅O: 382.14 (MH⁺).

[0911] 3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazine-2-yl)-N[4-chloro-2-fluorophenyl)methyl]benzamide: 3-(5-Amino-6-cyano-pyrazin-2-yl)-N-(4-chloro-2-fluoro-benzyl)-benzamide (238.0 mg, 0.63 mmol) and 1,1-dimethylethyl (3S)-3-(hydrazinocarbonyl)piperidine-1-carboxylate (460 mg, 1.89 mmol) was combined together and it was heated to 180 °C for 1 or 2 hrs. The reaction mixture was poured into water (575 mL), and extracted with ethyl acetate (2 \times 35 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by MPLC to give 3-[5-amino-6-(5-piperidin-3-yl-2H-[1,2,4]triazol-3-yl)-pyrazin-2-yl]-N-(4-chloro-2-fluoro-benzyl)-benzamide (206.0 mg, 65.0% yield): ¹H NMR (400 MHz, d₄-MeOH): δ 8.60 (d, 2H), 8.20 (d, 1H), 7.82 (d, 1H), 7.58 (t, 1H), 7.38 (t, 1H), 7.15 (m, 2H), 4.60(s, 2H), 3.62 (m, 1H), 3.40(m, 3H), 2.99 (m, 1H), 2.37 (m, 1H), 1.98 (m, 3H); MS (EI) for C₂₅H₂₄N₈OFCI: 507.03 (MH⁺).

[0912] 1-(1,1-dimethylethyl) 3-ethyl (3S)-piperidine-1,3-dicarboxylate: To an ice-cooled, stirred solution of (S) – (+) – ethyl nipecotate (3.95 g, 25.1 mmol, from Sigma-Aldrich) in 20 mL of THF, was added 8.21 g (37.6 mmol, 1.5 eq.) of di-*tert*-butyl dicarbonate. The ice bath was removed, and the reaction stirred at room temperature for 19 h and concentrated. The crude oil was purified by silica gel chromatography (55 mm column, 5 inches of silica gel) using 15% EtOAc in hexanes as eluent. Concentration afforded 6.28 g (97.2% yield) of product as an oil.

[0913] 1,1-dimethylethyl (3S)-3-(hydrazinocarbonyl)piperidine-1-carboxylate: To a solution of 6.28 g (24.4 mmol) of 1-(1,1-dimethylethyl) 3-ethyl (3S)-piperidine-1,3-dicarboxylate in 25 mL of MeOH was added 7.11 mL of hydrazine monohydrate (146 mmol, 5.98 eq.). The mixture was refluxed for 2 h and concentrated. The reaction could be monitored by LC/MS on a reversed-phase column at 220 nm wavelength. The crude material was taken up in EtOAc, and washed with 2 \times H₂O, 1 \times sat. aqueous NaCl, and dried over Na₂SO₄. Filtration and concentration afforded 3.21 g of product (54.0% yield)

as a viscous oil. The product was ca. 80% pure by hplc analysis and was used as such without further purification.

[0914] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.

[0915] 3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(2,6-difluorophenyl)methyl]benzamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.60 (d, 2H), 8.38 (d, 1H), 7.88 (d, 1H), 7.60 (t, 1H), 7.38 (m, 2H), 7.15 (t, 1H), 4.80 (s, 2H), 3.78 (m, 1H), 3.50 (m, 3H), 3.15 (m, 1H), 2.37 (m, 1H), 1.98 (m, 3H); MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_8\text{OF}_2$: 491.08 (MH^+).

[0916] 3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(2-chloro-6-fluorophenyl)methyl]benzamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.60 (d, 2H), 8.32 (d, 1H), 7.82 (d, 1H), 7.60 (t, 1H), 7.40 (t, 1H), 7.00 (m, 2H), 4.70 (s, 2H), 3.78 (m, 1H), 3.50 (m, 3H), 3.05 (m, 1H), 2.37 (m, 1H), 2.00 (m, 3H); MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_8\text{OFCl}$: 507.03 (MH^+).

[0917] 3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(2,5-dichlorophenyl)methyl]benzamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.75 (d, 2H), 8.38 (d, 1H), 7.95 (d, 1H), 7.62 (t, 1H), 7.40 (m, 2H), 7.25 (d, 1H), 4.62 (s, 2H), 3.78 (m, 1H), 3.50 (m, 3H), 3.05 (m, 1H), 2.41 (m, 1H), 2.00 (m, 3H); MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_8\text{OCl}_2$: 523.19 (M^+).

[0918] 3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(3,4-dichlorophenyl)methyl]benzamide: ^1H NMR (400 MHz, $\text{d}_4\text{-MeOH}$): δ 8.70 (d, 2H), 8.38 (d, 1H), 7.95 (d, 1H), 7.62 (t, 1H), 7.58 (s, 1H), 7.42 (d, 2H), 7.38 (d, 1H), 4.60 (s, 2H), 3.72 (m, 1H), 3.50 (m, 3H), 3.05 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_8\text{OCl}_2$: 523.19 (M^+).

(3-{5-Amino-6-[3-(2-thienyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol

[0919] To a solution of 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carbonitrile (149 mg, 0.74 mmol) was added thiophene-2-carboxylic acid hydrazide (94 mg, 0.66 mmol), sodium methoxide (53 mg, 0.996 mmol) in MeOH (5 mL). After reflux for 4 days, the reaction mixture was extracted with ethyl acetate (10 mL x 3) and dried over MgSO_4 . Filtration and concentration in vacuo gave a crude product, which was purified by preparative HPLC (0.1% TFA in water/0.1% TFA in CH_3CN). The product fractions

were neutralized by aqueous saturated sodium bicarbonate solution and partitioned with ethyl acetate. The aqueous phase was extracted once with additional ethyl acetate. The combined organic layers were washed with saturated aqueous sodium chloride then dried over anhydrous magnesium sulfate. Filtration and concentration in vacuo gave the (3-{5-amino-6-[3-(2-thienyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol (14 mg, 5 % yield): ^1H NMR (400 MHz, CD_3OD): δ 8.65 (s, 1H), 8.14 (s, 1H), 8.02 (d, 1H), 7.77 (m, 1H), 7.53 (m, 1H), 7.47 (m, 1H), 7.38 (m, 1H), 7.17 (m, 1H), 4.72 (s, 2H); MS (EI) for $\text{C}_{17}\text{H}_{14}\text{N}_6\text{O}$: 351.10 (MH^+).

[0920] (3-{5-Amino-6-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol: To a solution of 3-amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carbonitrile (250 mg, 1.11 mmol) was added 2-aminoisonicotinic acid hydrazide (311 mg, 2.04 mmol). After being heated at 180 °C for 90 min, the reaction mixture was dissolved in DMF and washed with water, ethyl acetate (10 mL x 3). The aqueous phase was extracted once with additional ethyl acetate. The combined organic layers were washed with saturated aqueous sodium chloride then dried over anhydrous magnesium sulfate. Filtration and concentration in vacuo gave a crude oil, which was purified by preparative HPLC to yield (3-{5-amino-6-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol (23.2 mg, 5.7 % yield): ^1H NMR (400 MHz, $\text{DMSO}-d_6$): δ 8.80 (s, 1H), 8.15 (s, 1H), 8.10 (m, 1H), 7.70 (m, 2H), 7.43 (m, 2H), 7.20 (m, 1H), 6.10 (s, 2H), 5.25 (m, 1H) 4.60 (m, 1H); MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_8\text{O}$: 361.11 (MH^+).

[0921] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following 1H-1,2,4-triazole compounds of the invention were prepared:

[0922] 5-{3-[(2,3-Dihydro-1H-inden-1-ylamino)methyl]phenyl}-3-{3-[3-(methyloxy)phenyl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine: ^1H NMR (400 MHz, $\text{DMSO}-d_6$): δ 8.82 (s, 1H), 8.38 (m, 2H), 7.82 (m, 1H), 7.76 (d, 1H), 7.62 (m, 2H), 7.52 (m, 1H), 7.43 (m, 1H), 7.35 (m, 3H), 7.11 (m, 1H), 4.75 (bs, 1H), 4.23 (m, 2H), 3.87 (s, 3H), 3.18 (m, 1H), 2.90 (m, 1H), 2.45 (m, 1H), 2.30 (m, 1H); MS (EI) for $\text{C}_{29}\text{H}_{27}\text{N}_7\text{O}$: 490.14 (MH^+).

[0923] 3-{5-Amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}-N-[(2-chloro-6-fluorophenyl)methyl]benzamide: ^1H NMR (400 MHz, $\text{DMSO}-d_6$): δ 10.40 (s, 1H), 8.97 (m, 2H), 8.57 (s, 1H), 8.40 (m, 1H), 7.87 (m, 1H), 7.80 (bs, 1H), 7.55 (m,

1H), 7.37 (m, 2H), 7.22 (m, 1H), 4.63 (s, 2H), 3.78 (m, 1H), 3.43 (m, 2H), 3.23 (m, 1H), 3.00 (m, 1H), 2.97 (s, 3H), 2.22 (m, 1H), 2.01 (m, 2H), 1.61 (m, 1H); MS (EI) for $C_{26}H_{26}N_8OFCl$: 521.02 (MH^+).

[0924] 3-(5-Amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(4-chloro-2-fluorophenyl)methyl]benzamide: 1H NMR (400 MHz, CD_3OD): δ 8.68 (s, 2H), 8.29 (d, 1H), 7.93 (d, 1H), 7.62 (t, 1H), 7.45 (t, 1H), 7.20 (m, 2H), 4.67 (s, 2H), 3.75 (m, 1H), 3.47 (m, 3H), 3.12 (m, 2H), 2.33 (m, 1H), 2.03 (m, 2H); MS (EI) for $C_{25}H_{24}N_8OFCl$: 507.02 (MH^+).

[0925] 3-(5-Amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(2,4-difluorophenyl)methyl]benzamide: 1H NMR (400 MHz, $DMSO-d_6$): δ 9.23 (m, 1H), 8.89 (m, 2H), 8.65 (s, 1H), 8.42 (m, 1H), 7.89 (m, 1H), 7.78 (bs, 1H), 7.60 (m, 1H), 7.45 (m, 1H), 7.26 (m, 1H), 7.08 (m, 1H), 4.54 (d, 2H), 3.30 (m, 2H), 3.18 (m, 1H), 2.96 (m, 1H), 2.19 (m, 1H), 1.87 (m, 3H); MS (EI) for $C_{25}H_{24}N_8OF_2$: 491.08 (MH^+).

[0926] 3-(5-Amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)-N-[(3,4-difluorophenyl)methyl]benzamide: 1H NMR (400 MHz, $DMSO-d_6$): δ 9.42 (m, 1H), 9.02 (m, 2H), 8.90 (s, 1H), 8.71 (s, 1H), 8.41 (m, 1H), 7.90 (m, 1H), 7.78 (m, 1H), 7.60 (m, 1H), 7.42 (m, 2H), 7.23 (m, 1H), 4.51 (d, 2H), 3.57 (m, 1H), 3.29 (m, 2H), 3.20 (m, 1H), 2.90 (m, 1H), 2.20 (m, 1H), 1.85 (m, 3H); MS (EI) for $C_{25}H_{24}N_8OF_2$: 491.08 (MH^+).

[0927] 3-[3-(2-Aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]-5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)pyrazin-2-amine: 1H NMR (400 MHz, $DMSO-d_6$): δ 8.99 (s, 1H), 8.85 (s, 1H), 7.31 (d, 2H), 8.07 (d, 1H), 7.85 (d, 1H), 7.73 (s, 1H), 7.55 (m, 2H), 7.37 (d, 1H), 7.31 (m, 1H), 4.83 (m, 1H), 4.32 (m, 2H), 3.25 (m, 1H), 2.91 (m, 1H), 2.43 (m, 1H); MS (EI) for $C_{27}H_{25}N_9$: 476.07 (MH^+).

[0928] (3-{5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol: 1H NMR (400 MHz, d_6 -DMSO): 10.34 (br s, 1H), 8.77 (s, 1H), 8.11 (s, 1H), 8.07 (d, 1H), 7.68 (br s, 2H), 7.42 (m, 1H), 7.34 (d, 1H), 4.58 (s, 2H), 3.75 (m, 1H), 3.45 (m, 1H), 3.38 (m, 1H), 3.22 (m, 1H), 2.99 (m, 1H), 2.82 (m, 3H), 2.19 (m, 1H), 1.95 (m, 2H), 1.68 (m, 1H); MS (EI) for $C_{19}H_{23}N_7O$: 366.2 (MH^+).

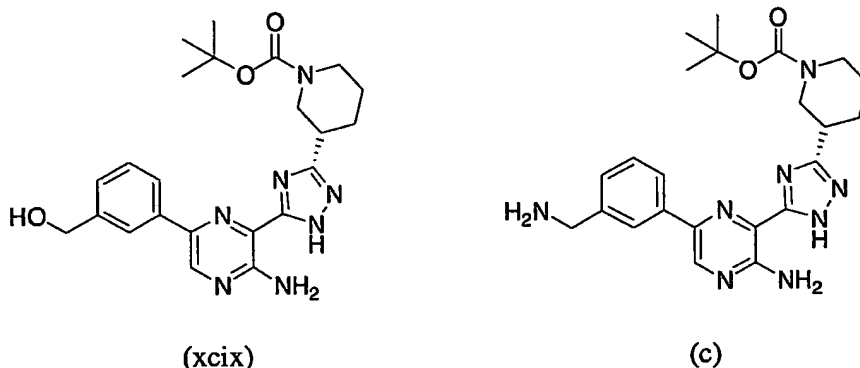
[0929] 5-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-3-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-amine: 1H NMR (400 MHz, d_6 -DMSO): 10.40 (br s, 1H), 9.76 (br s, 1H), 8.85 (s, 1H), 8.67 (s, 1H), 8.25 (m, 1H), 7.82

(d, 1H), 7.75 (br s, 2H), 7.53 (m, 2H), 7.36 (m, 2H), 7.30 (m, 1H), 4.83 (m, 1H), 4.27 (m, 2H), 3.76 (m, 1H), 3.46 (m, 1H), 3.40 (m, 1H), 3.20 (m, 2H), 2.94 (m, 2H), 2.82 (m, 3H), 2.38 (m, 1H), 2.20 (m, 1H), 1.96 (m, 2H), 1.67 (m, 1H); MS (EI) for $C_{28}H_{32}N_8$: 481.2 (MH^+).

Example 59

[0930] Scheme 23 depicts two exemplary compounds, (xcix) and (c), according to formula I, which are useful as intermediates to synthesize other compounds of the invention. Exemplary syntheses of such compounds follows, and generally can also be used to make other heterocycles (besides triazoles) for A, according to formula I.

Scheme 23



[0931] 1,1-Dimethylethyl (3R)-3-(5-{3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl}-1H-1,2,4-triazol-3-yl)piperidine-1-carboxylate: 3-Amino-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carbonitrile (514.0 mg, 2.27 mmol) and 1,1-dimethylethyl (3S)-3-(hydrazinocarbonyl)piperidine-1-carboxylate (1.66g, 6.83 mmol) were combined together and heated to 180 °C for 1 - 2 h. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 35 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by MPLC to give 1,1-dimethylethyl (3R)-3-(5-{3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl}-1H-1,2,4-triazol-3-yl)piperidine-1-carboxylate. (655.0 mg, 64.2 % yield): MS (EI) for $C_{23}H_{29}N_7O_3$: 452.29 (MH^+).

[0932] 1,1-dimethylethyl(3R)-3-(5-{3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl}-1H-1,2,4-triazol-3-yl)piperidine-1-carboxylate: 1,1-dimethylethyl (3R)-3-(5-{3-amino-6-

[3-(hydroxymethyl)phenyl]pyrazin-2-yl}-1H-1,2,4-triazol-3-yl)piperidine-1-carboxylate (655.0 mg, 1.45 mmol) was dissolved in THF (20 mL) followed by the addition of diphenylphosphoryl azide (0.45 mL, 2.03 mmol) and DBU (0.31 mL, 2.03 mmol). It was stirred at room temperature overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography (50% Ethylacetate/hexanes) to give the intermediate azide, 458.0 mg (66.4 %, MS (EI) for $C_{23}H_{30}N_8O_2$: 451.27 (MH⁺), which was dissolved in the mixture of THF (15 mL) and H₂O (1.5 mL) followed by the addition of triphenylphosphine (0.76g, 2.88 mmol) and it was stirred at room temperature overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate. The extracts were filtered, concentrated, and purified by column chromatography (0 to 100% MeOH in Ethyl acetate) to give 1,1-dimethylethyl(3R)-3-(5-{3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl}-1H-1,2,4-triazol-3-yl)piperidine-1-carboxylate, 368.0 mg (85.0 % yield).

[0933] 5-[3-(Aminomethyl)phenyl]-3-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine: To a solution of 1,1-dimethylethyl(3R)-3-(5-{3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl}-1H-1,2,4-triazol-3-yl)piperidine-1-carboxylate (50.0 mg, 0.11 mmol) in MeOH (3.0 mL) was added 4.0M HCl in dioxane (15 mL) and stirred it overnight at room temperature. All solvent was evaporated under vacuum to give compound 1, 28.0 mg (72.0 % yield): ¹H NMR (400 MHz, d₄-MeOH): δ 8.62 (s, 1H), 8.38 (s, 1H), 8.21 (d, 1H), 7.60 (m, 1H), 4.22(s, 2H), 3.65 (m, 1H), 3.42 (m, 3H), 3.16 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $C_{18}H_{22}N_8$: 351.27 (MH⁺).

Example 60

[0934] Generally, intermediates (xcix) and (c) (see Scheme 23 above) are used to attach Y (according to formula I) to make compounds of the invention, and thereby also establish X in many examples. Included in this example is description of such syntheses.

[0935] N-([3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl)methyl]-2-chlorobenzamide HCl: Intermediate 2 (50.0 mg, 0.11 mmol) was dissolved in DMF (5mL) followed by the addition of HOBt (30.0 mg, 0.22 mmol) and EDCI (43.0 mg, 0.22 mmol). It was stirred at room temperature overnight. The reaction

mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate and filtered, concentrated and purified by column chromatography (50% Ethylacetate/hexanes) and subsequently it was treated with 4.0 M HCl in dioxane (20 mL) to give compound 1, 42.2 mg (78.0 % yield). ¹H NMR (400 MHz, d4-MeOH): δ 8.50 (s, 1H), 8.20 (s, 1H), 8.11 (s, 1H), 7.58 (m, 6H), 4.67(s, 2H), 3.73 (m, 1H), 3.45 (m, 3H), 3.18 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for C₂₅H₂₅N₈OCl: 489.24 (MH⁺).

[0936] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.

[0937] N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl)methyl]-4-chlorobenzamide: ¹H NMR (400 MHz, d4-MeOH): δ 8.52 (s, 1H), 8.14 (s, 1H), 8.00 (d, 1H), 7.84 (d, 2H), 7.42 (m, 4H), 4.62(s, 2H), 3.65 (m, 2H), 3.42 (m, 2H), 3.16 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for C₂₅H₂₅N₈OCl: 489.20 (MH⁺).

[0938] N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl)methyl]-4-iodobenzamide: ¹H NMR (400 MHz, d4-MeOH): δ 8.58 (s, 1H), 8.18 (s, 1H), 7.98 (d, 1H), 7.81 (d, 2H), 7.62 (m, 2H), 7.40 (m, 2H), 4.62(s, 2H), 3.64 (m, 2H), 3.42 (m, 2H), 3.17 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for C₂₅H₂₅N₈OI: 581.16 (MH⁺).

[0939] N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl)methyl]-3,5-difluorobenzamide: ¹H NMR (400 MHz, d4-MeOH): δ 8.55 (s, 1H), 8.18 (s, 1H), 7.98 (d, 1H), 7.42 (m, 4H), 7.18 (m, 1H), 4.62(s, 2H), 3.70 (m, 2H), 3.42 (m, 2H), 3.17 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for C₂₅H₂₄N₈OF₂: 491.21 (MH⁺).

[0940] N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl)methyl]-4-bromo-2-fluorobenzamide: ¹H NMR (400 MHz, d4-MeOH): δ 8.50 (s, 1H), 8.18 (s, 1H), 8.00 (d, 1H), 7.62 (t, 1H), 7.42 (m, 4H), 4.62(s, 2H), 3.63 (m, 2H), 3.42 (m, 2H), 3.17 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for C₂₅H₂₄N₈OFBr: 553.16 (M+2).

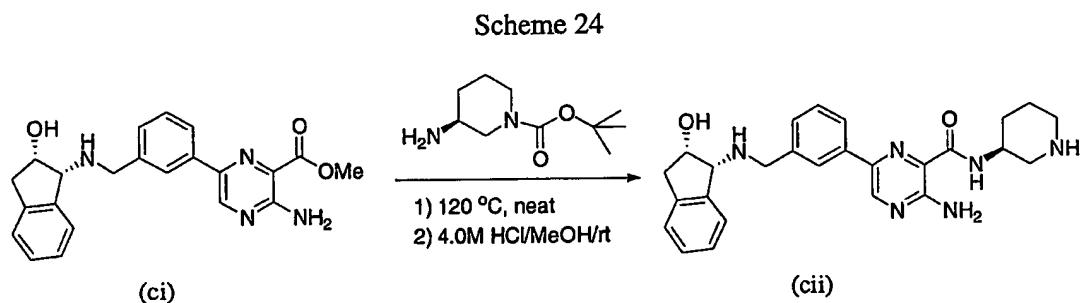
[0941] N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl)methyl]-4-bromo-2-chlorobenzamide: ¹H NMR (400 MHz, d4-MeOH): δ 8.50 (s, 1H), 8.18 (s, 1H), 8.00 (d, 1H), 7.62 (s, 1H), 7.42 (m, 4H), 4.62(s, 2H), 3.63 (m, 2H),

3.42 (m, 2H), 3.17 (m, 1H), 2.38 (m, 1H), 2.00 (m, 3H); MS (EI) for $C_{25}H_{24}N_8OClBr$: 567.14 (M^+).

[0942] 3-amino-6-{3-[[[(4-chlorophenyl)amino]carbonyl]amino)methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.42 (s, 1H), 9.30 (m, 1H), 9.12 (m, 1H), 8.71 (d, 1H), 8.15 (s, 1H), 8.00 (d, 1H), 7.42 (m, 3H), 7.38 (m, 3H), 4.80 (br, 2H), 4.40 (s, 2H), 4.35 (m, 1H), 3.60 (m, 1H), 3.20 (m, 3H), 2.72 (m, 1H), 1.80 (m, 4H); MS (EI) for $C_{24}H_{26}N_7O_2Cl$: 480.19 (MH^+).

Example 61

[0943] Scheme 24 depicts synthesis of amides of the invention according to formula I, from corresponding esters. In general, esters, for example pyrazine ester (ci), are combined with amines, for example as depicted, to give amides, for example (cii), according to formula I.



3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

[0944] A mixture of 3-amino-6-[3-[(2-hydroxy-indan-1-ylamino)-methyl]-phenyl]-pyrazine-2-carboxylic acid methyl ester (400.0 mg, 1.02 mmol) and 1,1-dimethylethyl (3S)-3-aminopiperidine-1-carboxylate (970.0 mg, 4.84 mmol) was heated to 120 °C overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2 × 50 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography (3% MeOH/Dichloromethane) and subsequently it was treated with 4.0 M HCl in dioxane (20 mL) to give 3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide, 275.0 mg (58.7 % yield). 1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 2H), 8.18 (s, 1H), 7.80 (d, 1H), 7.60 (m, 2H), 7.38 (m, 3H), 4.80 (s,

1H), 4.50(s, 2H), 4.40 (m, 1H), 3.65 (m, 2H), 3.42 (m, 5H), 3.16 (m, 2H), 2.15 (m, 2H), 1.87 (m, 1H); MS (EI) for C₂₆H₃₀N₆O₂: 459.31 (MH⁺).

Example 62

3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-ethylpyrazine-2-carboxamide

[0945] To a solution of 3-amino-*N*-ethyl-6-[3-(hydroxymethyl)phenyl]pyrazine-2-carboxamide (566.1 mg, 1.0 eq) in dichloromethane (20 mL) were added carbon tetrabromide (749.8 mg, 1.1eq), triphenylphosphine (600 mg, 1.1 eq) at ambient temperature. After being stirred for 7 h, the reaction mixture was partitioned with dichloromethane (10 mL x 3), washed with brine. The organic layer was combined and dried over anhydrous magnesium sulfate. Filtration and concentration in vacuo gave a crude product. Purification by column chromatography afforded the 3-amino-6-[3-(bromomethyl)phenyl]-*N*-ethylpyrazine-2-carboxamide (534 mg, 77 % yield): ¹H NMR (400 MHz, DMSO-d₆): δ 8.87 (m, 2H), 8.84 (s, 2H), 8.23 (m, 1H), 7.61 (bs, 1H), 7.45 (m, 1H), 4.77 (s, 2H), 3.38 (m, 2H), 1.16 (m, 3H).

[0946] The 3-amino-6-[3-(bromomethyl)phenyl]-*N*-ethylpyrazine-2-carboxamide (112mg, 1.0eq) was dissolved in acetonitrile (5 mL) and treated with (1*S*)-2,3-dihydro-1*H*-inden-1-amine (133.8mg, 3.0eq), diisopropylethylamine (69.3mg, 1.6 eq) at ambient temperature. After being stirred for 7h, the white suspension was filtered on Celite and concentrated in vacuo. The resulting residue was purified by flash column chromatography to give the 3-amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)-*N*-ethylpyrazine-2-carboxamide (50.2mg, 39 % yield): ¹H NMR (400 MHz, CD₃OD): δ 8.72 (s, 1H), 8.21 (s, 1H), 8.11 (d, 1H), 7.61 (d, 1H), 7.58 (t, 1H), 7.49 (d, 1H), 7.39 (d, 2H), 7.39 (d, 2H), 7.37 (m, 1H), 4.38 (s, 2H), 3.47 (m 1H), 3.23 (m, 1H), 3.01 (m, 1H), 2.64 (m, 1H), 2.39 (m, 1H); MS (EI) for C₂₃H₂₅N₅O: 388.22 (MH⁺).

[0947] 3-Amino-6-(3-([(5,7-difluoro-2,3-dihydro-1*H*-inden-1-yl)amino]methyl)phenyl)-*N*-ethylpyrazine-2-carboxamide: The 3-Amino-6-[3-(bromomethyl)phenyl]-*N*-ethylpyrazine-2-carboxamide (100mg, 1.0eq) was dissolved in acetonitrile (5 mL) and treated with [(1*S*)-5,7-difluoro-2,3-dihydro-1*H*-inden-1-yl]amine (151mg, 3.0eq), diisopropylethylamine (62mg, 1.6eq) at ambient temperature. After being stirred for 8h, the white suspension was concentrated in vacuo and purified by flash column

chromatography to give the title compound as colorless oil, which was dissolved in ethyl acetate (10 mL) and treated 1M HCl in ether. The resulting precipitate was filtered, washed with ether (2 x 2mL), and dried to afford 42mg (12 % yield) of 3-amino-6-(3-[[[5,7-difluoro-2,3-dihydro-1H-inden-1-yl]amino]methyl]phenyl)-N-ethylpyrazine-2-carboxamide hydrochloride as a yellow solid: ^1H NMR (400 MHz, CD_3OD): δ 8.71 (s, 1H), 8.22 (s, 1H), 8.13 (d, 1H), 7.57 (t, 1H), 7.51 (m, 1H), 7.06 (m, 1H), 6.98 (m, 1H), 5.08 (m, 1H), 4.38 (q, 2H), 3.45 (m, 2H), 3.39 (m, 1H), 3.05 (m, 1H), 2.84 (m, 1H), 2.53 (m, 1H); MS (EI) for $\text{C}_{23}\text{H}_{23}\text{N}_5\text{OF}_2$: 424.16 (MH^+).

[0948] 3-Amino-6-(3-[[[1S]-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl)-N-(naphthalen-2-ylmethyl)pyrazine-2-carboxamide: To a solution of methyl 3-amino-6-(3-[[[1S]-2,3-dihydro-1H-inden-1-ylaminomethyl]phenyl]pyrazine-2-carboxylate (1.0eq) in methyl alcohol was added 2-aminonaphthalene (10.0eq) and the reaction mixture was allowed to heat at 85°C overnight. The solvent was removed followed by preparative HPLC (0.1% TFA in water/0.1% TFA in CH_3CN), which was neutralized by aqueous saturated sodium bicarbonate solution and partitioned with ethyl acetate. The aqueous phase was extracted once with additional ethyl acetate. The combined organic layers were washed with saturated aqueous sodium chloride then dried over anhydrous magnesium sulfate. Filtration and concentration to ca. 1 mL followed by addition of 1.0 M HCl in ether. The resulting precipitate was filtered and dried to give 3-amino-6-(3-[[[1S]-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl)-N-(naphthalen-2-ylmethyl)pyrazine-2-carboxamide hydrochloride as a yellow solid: ^1H NMR (400 MHz, CD_3OD): δ 8.66 (s, 1H), 8.10 (d, 1H), 8.01 (m, 2H), 7.82 (d, 1H), 7.78 (d, 1H), 7.44 (m, 4H), 7.39 (m, 2H), 7.30 (m, 2H), 7.21 (m, 1H), 5.04 (s, 2H), 4.79 (m, 1H), 4.21 (s, 2H), 3.12 (m, 1H), 2.90 (m, 1H), 2.51 (m, 1H), 2.20 (m, 1H); MS (EI) for $\text{C}_{32}\text{H}_{29}\text{N}_5\text{O}$: 501.17 (MH^+).

[0949] 3-Amino-N-cyclohexyl-6-(3-[[[1S]-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, CD_3OD): δ 8.70 (s, 1H), 8.11 (m, 2H), 7.62 (m, 2H), 7.53 (m, 1H), 7.41 (m, 2H), 7.33 (m, 1H), 4.40 (s, 2H), 3.93 (m, 1H), 3.15 (m, 1H), 2.75 (m, 1H), 2.41 (m, 1H), 2.00 (m, 2H), 1.84 (m, 2H), 1.75 (m, 1H), 1.50 (m, 2H); MS (EI) for $\text{C}_{27}\text{H}_{31}\text{N}_5\text{O}$: 442.24 (MH^+).

[0950] 3-Amino-6-(3-[[[1S]-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl)-N-(pyridin-4-ylmethyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, CD_3OD): δ 8.78 (m, 3H), 8.70 (m, 1H), 8.14 (m, 3H), 7.74 (d, 1H), 7.57 (t, 1H), 7.50 (m, 1H), 7.38 (m, 2H),

7.22 (m, 1H), 4.92 (s, 2H), 4.40 (s, 2H), 3.21 (m, 1H), 3.00 (m, 1H), 2.61 (m, 1H), 2.42 (m, 1H); MS (EI) for $C_{27}H_{31}N_5O$: 451.12 (MH^+).

[0951] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-pyrrolidin-1-ylethyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.83 (s, 1H), 8.78 (s, 1H), 8.15 (d, 1H), 7.91 (d, 1H), 7.56 (m, 2H), 7.41 (m, 2H), 7.38 (m, 1H), 4.41 (s, 2H), 3.82 (m, 4H), 3.54 (m, 2H), 3.24 (m, 1H), 3.18 (m, 1H), 3.05 (m, 1H), 2.83 (m, 1H), 2.43 (m, 1H), 2.17 (m, 2H), 2.10 (m, 2H); MS (EI) for $C_{27}H_{32}N_6O$: 457.21 (MH^+).

[0952] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-piperidin-1-ylethyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.88 (t, 1H), 8.76 (s, 1H), 8.09 (m, 1H), 7.79 (d, 1H), 7.54 (m, 2H), 7.38 (m, 2H), 7.31 (m, 1H), 4.92 (m, 1H), 4.42 (s, 1H), 4.4 – 3.8 (m, 4H), 3.83 (t, 1H), 3.72 (d, 1H), 3.44 (t, 1H), 3.40 (m, 1H), 3.05 (m, 2H), 2.66 (m, 1H), 2.52 (m, 1H), 1.80 (m, 4H), 1.49 (m, 1); MS (EI) for $C_{28}H_{34}N_6O$: 471.20 (MH^+).

[0953] 3-amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-morpholin-4-ylethyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.90 (s, 1H), 8.75 (s, 1H), 8.08 (m, 1H), 7.78 (d, 1H), 7.57 (d, 2H), 7.38 (m, 2H), 7.32 (m, 1H), 4.43 (s, 2H), 4.25 (m, 3H), 3.94 (t, 2H), 3.87 (m, 2H), 3.77 (m, 2H), 3.57 (t, 2H), 3.32 (m, 2H), 3.04 (m, 1H), 2.66 (m, 1H), 2.51 (m, 1H); MS (EI) for $C_{27}H_{32}N_6O_2$: 471.20 (MH^+).

[0954] 3-amino-N-(cyclopropylmethyl)-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.58 (s, 1H), 8.13 (t, 1H), 7.91 (s, 1H), 7.72 (m, 1H), 7.40 (m, 3H), 7.17 (m, 2H), 4.32 (t, 1H), 3.45 (s, 2H), 3.33 (t, 2H), 3.02 (m, 1H), 2.72 (m, 1H), 2.46 (m, 1H), 1.90 (m, 1H), 1.20 (m, 2H), 0.55 (m, 2H), 0.37 (m, 2H); MS (EI) for $C_{25}H_{27}N_5O$: 414.21 (MH^+).

[0955] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(phenylmethyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.70 (s, 1H), 8.39 (m, 1H), 8.14 (m, 1H), 7.66 (d, 1H), 7.54 (t, 1H), 7.52 (m, 1H), 7.40 (m, 3H), 7.11 (m, 1H), 7.24 (m, 1H), 4.66 (s, 2H), 4.39 (s, 2H), 3.23 (m, 1H), 3.02 (m, 1H), 2.61 (m, 1H), 2.41 (m, 1H); MS (EI) for $C_{28}H_{27}N_5O$: 451.20 (MH^+).

[0956] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(2-phenylethyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.68 (s, 1H), 8.35 (m, 1H), 8.08 (m, 1H), 7.69 (d, 1H), 7.57 (m, 2H), 7.40 (m, 1H), 7.32 (m, 6H), 7.21

(m, 1H), 4.95 (m, 1H), 4.39 (s, 2H), 3.64 (q, 2H), 3.23 (m, 1H), 3.06 (m, 1H), 3.92 (t, 2H), 2.65 (m, 1H), 2.42 (m, 1H); MS (EI) for $C_{29}H_{29}N_5O$: 464.20 (MH^+).

[0957] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[2-(dimethylamino)ethyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.708 (s, 1H), 8.74 (m, 1H), 8.13 (m, 1H), 7.58 (d, 1H), 7.59 (t, 1H), 7.48 (m, 1H), 7.40 (m, 2H), 7.35 (m, 1H), 4.40 (s, 2H), 3.78 (t, 2H), 3.50 (t, 2H), 3.59 (m, 1H), 3.02 (m, 7H), 2.65 (m, 1H), 2.45 (m, 1H); MS (EI) for $C_{25}H_{30}N_6O$: 431.22 (MH^+).

[0958] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.81 (s, 1H), 8.65 (s, 1H), 8.63 (s, 1H), 8.10 (d, 1H), 7.73 (d, 1H), 7.56 (m, 2H), 7.41 (m, 2H), 7.38 (m, 1H), 4.41 (m, 2H), 3.80 (t, 2H), 3.28 (m, 1H), 3.17 (t, 2H), 3.04 (m, 1H), 2.65 (m, 1H), 2.57 (m, 1H), 2.17 (m, 2H); MS (EI) for $C_{26}H_{27}N_7O$: 454.18 (MH^+).

[0959] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[2-(methylamino)ethyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.80 (s, 1H), 8.78 (s, 1H), 8.12 (d, 1H), 7.79 (d, 1H), 7.57 (m, 2H), 7.39 (m, 2H), 7.33 (m, 1H), 4.40 (s, 2H), 3.82 (t, 2H), 3.27 (m, 1H), 3.17 (t, 2H), 3.02 (m, 1H), 2.67 (m, 1H), 2.43 (m, 1H); MS (EI) for $C_{24}H_{28}N_6O$: 417.21 (MH^+).

[0960] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[3-(dimethylamino)propyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.80 (s, 1H), 8.72 (s, 1H), 8.11 (d, 1H), 7.79 (d, 1H), 7.58 (m, 2H), 7.39 (m, 2H), 7.32 (m, 1H), 4.40 (s, 2H), 3.61 (t, 2H), 3.52 (m, 5H), 3.05 (m, 1H), 2.71 (m, 1H), 2.49 (m, 1H), 2.12 (m, 2H); MS (EI) for $C_{26}H_{32}N_6O$: 445.21 (MH^+).

[0961] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[3-(1H-imidazol-1-yl)propyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 9.10 (s, 1H), 8.68 (s, 1H), 8.03 (d, 2H), 7.51 (m, 2H), 7.46 (m, 3H), 7.30 (d, 1H), 7.22 (m, 2H), 4.31 (m, 4H), 3.48 (t, 2H), 3.41 (m, 1H), 2.93 (m, 1H), 2.52 (m, 1H), 2.38 (m, 1H), 2.23 (m, 1H); MS (EI) for $C_{27}H_{29}N_7O$: 468.18 (MH^+).

[0962] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-N-[3-(methylamino)propyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, CD_3OD): δ 8.64 (s, 1H), 8.63 (s, 1H), 8.02 (d, 1H), 7.43 (m, 2H), 7.31 (m, 2H), 7.22 (m, 1H), 4.85 (m, 1H), 4.31 (s, 2H), 3.50 (t, 2H), 3.18 (m, 1H), 2.95 (m, 3H), 2.62 (s, 2H), 2.53 (m, 1H), 2.37 (m, 1H), 1.98 (m, 1H); MS (EI) for $C_{25}H_{30}N_6O$: 431.22 (MH^+).

- [0963] 3-Amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)-*N*-(3-pyrrolidin-1-ylpropyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, CD₃OD): δ 8.83 (s, 1H), 8.73 (s, 1H), 8.10 (d, 1H), 7.79 (d, 1H), 7.59 (m, 2H), 7.38 (m, 2H), 7.32 (m, 1H), 4.43 (s, 2H), 3.67 (m, 4H), 3.20 (m, 4H), 2.63 (m, 2H), 2.48 (m, 1H), 2.14 (m, 1H), 2.02 (m, 2H); MS (EI) for C₂₈H₃₄N₆O: 471.22 (MH⁺).
- [0964] 3-Amino-*N*-1-azabicyclo[2.2.0]oct-3-yl-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, CD₃OD): δ 8.77 (s, 1H), 8.45 (m, 1H), 8.12 (d, 1H), 7.52 (d, 1H), 7.56 (t, 2H), 7.49 (d, 1H), 7.40 (d, 2H), 7.35 (m, 1H), 4.55 (m, 1H), 4.40 (s, 2H), 3.79 (d, 2H), 3.63 (d, 1H), 3.03 (m, 1H), 2.68 (m, 1H), 2.43 (m, 2H), 2.31 (m, 1H), 2.15 (m, 2H), 2.31 (m, 1H), 2.15 (m, 2H), 1.83 (m, 1H); MS (EI) for C₂₈H₃₂N₆O: 469.13 (MH⁺).
- [0965] 3-Amino-*N*-[(3*R*)1-azabicyclo[2.2.0]oct-3-yl]-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-*d*₆): δ 10.22 (bs, 1H), 10.10 (m, 1H), 9.95 (m, 1H), 9.08 (d, 1H), 8.93 (s, 2H), 8.12 (m, 1H), 7.91 (d, 1H), 7.70 (bs, 2H), 7.50 (d, 2H), 7.37 (d, 2H), 7.31 (m, 1H), 4.82 (bs, 1H), 4.49 (m, 1H), 4.38 (t, 2H), 4.02 (m, 1H), 3.60 (t, 1H), 3.21 (m, 4H), 2.90 (m, 1H), 2.44 (m, 2H), 2.13 (m, 1H), 1.94 (m, 2H), 1.73 (m, 1H); MS (EI) for C₂₈H₃₂N₆O: 469.15 (MH⁺).
- [0966] 3-Amino-*N*-azepan-3-yl-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-*d*₆): δ 9.94 (bs, 2H), 9.58 (bs, 1H), 9.33 (bs, 1H), 9.10 (d, 1H), 8.91 (m, 2H), 8.09 (m, 1H), 7.88 (m, 1H), 7.47 (m, 2H), 7.42 (m, 2H), 7.27 (m, 2H), 4.82 (m, 1H), 4.49 (m, 1H), 4.36 (m, 2H), 3.48 (m, 1H), 3.30 (m, 3H), 3.05 (m, 1H), 2.82 (m, 1H), 2.48 (m, 2H), 2.10 (m, 1H), 2.00 (m, 1H), 1.85 (m, 3H), 1.62 (m, 1H); MS (EI) for C₂₇H₃₂N₆O: 457.12 (MH⁺).
- [0967] 3-amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)pyrazine-2-carbohydrazide: ¹H NMR (400 MHz, *d*₆-DMSO): 9.96 (m, 1H), 8.80 (s, 1H), 8.16 (s, 1H), 8.01 (d, 1H), 7.56 (br s, 2H), 7.38 (m, 3H), 7.19 (m, 1H), 7.15 (m, 2H), 4.57 (d, 2H), 4.16 (m, 1H), 3.86 (m, 2H), 2.92 (m, 1H), 2.72 (m, 1H), 2.53 (m, 1H), 2.31 (m, 1H), 1.81 (m, 1H); MS (EI) for C₂₁H₂₂N₆O: 375.2 (MH⁺).
- [0968] 3-amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)-*N*-(5-hydroxypentyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, CDCl₃): 8.59 (s, 1H), 8.18 (t, 1H), 7.95 (s, 1H), 7.70 (d, 1H), 7.39 (m, 3H), 7.21 (m, 3H), 4.35 (t, 1H), 3.98 (m, 2H), 3.60 (m, 2H), 3.47 (m, 2H), 3.03 (m, 1H), 2.84 (m, 1H), 2.47 (m, 1H), 2.04 (br s, 3H),

1.93 (m, 1H), 1.68 (m, 2H), 1.60 (m, 2H), 1.51 (m, 2H); MS (EI) for $C_{26}H_{31}N_5O_2$: 446.2 (MH^+).

[0969] 3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N-(4-hydroxybutyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, $CDCl_3$): 8.84 (m, 1H), 8.60 (s, 1H), 8.23 (s, 1H), 7.69 (d, 1H), 7.36 (m, 2H), 7.28 (m, 1H), 7.24 (m, 1H), 7.21 (m, 2H), 4.37 (t, 1H), 3.96 (m, 2H), 3.47 (m, 4H), 3.05 (m, 1H), 2.85 (m, 1H), 2.63 (br s, 2H), 2.46 (m, 1H), 1.98 (m, 1H), 1.76 (m, 2H), 1.63 (m, 2H); MS (EI) for $C_{25}H_{29}N_5O_2$: 432.2 (MH^+).

[0970] 3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N'-methylpyrazine-2-carbohydrazide: 1H NMR (400 MHz, $CDCl_3$): 9.24 (s, 1H), 8.66 (s, 1H), 7.91 (s, 1H), 7.74 (m, 1H), 7.44 (m, 2H), 7.41 (m, 1H), 7.25 (m, 1H), 7.22 (m, 2H), 4.35 (t, 1H), 4.00 (m, 2H), 3.05 (m, 1H), 2.85 (m, 1H), 2.77 (s, 3H), 2.48 (m, 1H), 1.93 (m, 1H); MS (EI) for $C_{22}H_{24}N_6O$: 389.2 (MH^+).

[0971] 3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl}phenyl)-N,N-dimethylpyrazine-2-carboxamide: 1H NMR (400 MHz, $CDCl_3$): 8.51 (s, 1H), 7.89 (s, 1H), 7.74 (m, 1H), 7.40 (m, 3H), 7.22 (m, 3H), 5.86 (br s, 2H), 4.32 (t, 1H), 3.98 (m, 2H), 3.31 (s, 3H), 3.17 (s, 3H), 3.03 (m, 1H), 2.83 (m, 1H), 2.45 (m, 1H), 1.91 (m, 1H); MS (EI) for $C_{23}H_{25}N_5O$: 388.2 (MH^+).

Example 63

Synthesis of compounds according to formula I, wherein A is $-(C=O)-$ or $-(C=NR^6)-$ derived from methyl 3-amino-2-pyrazine carboxylate.

[0972] Methyl 3-amino-6-bromopyrazine-2-carboxylate. To a solution of methyl 3-amino-2-pyrazine carboxylate (30.0 g, 200 mmol) in acetic acid (200 mL), bromine (11 mL) was added slowly via addition funnel. After complete addition of bromine, sodium carbonate powder was added slowly until precipitation occurred. The reaction mixture was allowed to stir at room temperature for 2 h. The reaction mixture was reduced to half-volume in vacuo and then diluted with water (500 mL). The reaction mixture was shaken vigorously and the resulting solid was collected using vacuum filtration. The solid was washed with ether to afford a pure yellow solid (91 % yield).

[0973] 3-Amino-6-bromopyrazine-2-carboxylic acid. To a solution of methyl-3-amino-6-bromopyrazine carboxylate (10 g, 43 mmol) in THF:H₂O (3:1), lithium hydroxide (6.0 g,

140 mmol) was added. The reaction mixture was stirred at room temperature and monitored by LCMS. The reaction was complete after 1 h. The reaction mixture was poured into 1 N HCl (250 mL), extracted with ethyl acetate (3x, 200 mL), washed with brine (3x, 100 mL), dried over anhydrous sodium sulfate, filtered and concentrated to afford a yellow solid (100 % yield).

[0974] 3-Amino-6-(3-[[[(biphenyl-2-ylmethyl)amino]carbonyl}phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: To a solution of 3-amino-6-bromopyrazine-2-carboxylic acid (4.0 g, 18 mmol) in DMF, HATU (10.4 g, 27.5 mmol), *N,N*-diisopropylethylamine (8.5 mL, 27.5 mmol) and (*S*)-3-amino-1-*N*-Boc-piperidine (4.0 g, 20 mmol, commercially available from Astatech and Arch Corporation) were added. The reaction was allowed to stir at room temperature for 1 h. The reaction mixture was diluted with water (100 mL), extracted with ethyl acetate (3x, 200 mL), washed with 5 % LiCl (3x, 100 mL) and brine (3x, 100 mL), dried over anhydrous sodium sulfate, filtered and concentrated in vacuo to afford a brown solid (MS (ESI-LCMS) for $C_{15}H_{22}BrN_5O_3$: 400 (MH^+)). This material was taken up in ethylene glycol dimethyl ether (150 mL), and triethylamine (4.6 mL, 45 mmol), $Pd(dppf)_2Cl_2$ (2.3 g, 2.75 mmol), 3-carboxyphenylboronic acid (3.6 g, 21.8 mmol) and water (15 mL) were added. The reaction was refluxed at 85 °C and monitored by LCMS. The reaction was complete after 3 h. The solvent was removed under reduced pressure to afford a brown solid. The crude product was purified by silica gel chromatography to afford a yellow solid (MS (ESI-LCMS) for $C_{22}H_{27}N_5O_5$: 442 (MH^+)). A portion of this intermediate (50 mg, 0.11 mmol) was dissolved in DMF (1mL), and HATU (62.7 mg, 0.165 mmol), *N,N*-diisopropylethylamine (29 μ L, 0.16 mmol) and 2-phenylbenzylamine (24.2 mg, 0.132 mmol) were added. The reaction was stirred at room temperature and monitored by LCMS. The reaction was complete after 1 h. The crude product was purified by preparative HPLC and lyophilized to afford a white solid (MS (ESI-LCMS) for $C_{35}H_{38}N_6O_4$: 607 (MH^+)). This material was dissolved in methanol (2 mL), and 4M HCl in dioxane (2 mL) was added. The reaction was stirred at room temperature and monitored by LCMS. The reaction was complete after 1 h. The solvents were removed under reduced pressure to afford a yellow gel product. The yellow gel product was dissolved in acetonitrile-water (2 mL), frozen and lyophilized to afford a yellow solid: 1H NMR (400 MHz, d_6 -DMSO): δ 9.25 (t, 1H), 9.05 (d, 1H), 8.95 (s, 1H), 8.9 (d, 1H), 8.85 (d, 1H), 8.6 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.5-7.3 (m,

8H), 7.25 (m, 1H), 4.5 (d, 2H), 4.35 (m, 1H), 3.25 (m, 2H), 3.05 (m, 1H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{30}H_{30}N_6O_2$: 507 (MH⁺).

[0975] 3-Amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: To a solution of 3-carboxyphenyl boronic acid (20.0 g, 120 mmol) in thionyl chloride (120 mL) was added DMF (3 drops). The reaction mixture was allowed to stir at room temperature overnight. Excess thionyl chloride was removed under reduced pressure to afford a white solid. To a solution of this residue and triethylamine (33.4 mL, 240 mmol) in THF (200 mL) at 0 °C was added slowly (*S*)-(+)-1-aminoindane (19.2 g, 144 mmol) in MeOH (100 mL) via addition funnel. After addition of (*S*)-(+)-1-aminoindane, the reaction mixture was stirred at room temperature and monitored by LCMS. The reaction was complete after 3 h. The reaction mixture was then concentrated under reduced pressure to afford a white solid. This material (33.72 g, 120 mmol) was added to a solution of methyl-3-amino-6-bromopyrazine carboxylate (23.2 g, 100 mmol) in DMF (200 mL), and triethylamine (28 mL, 200 mmol), Pd(dppf)₂Cl₂ (8.16 g, 10 mmol) and water (20 mL) were added. The reaction mixture was stirred at 80 °C overnight. The reaction mixture was cooled to room temperature. Water (800 mL) was added and the precipitated solid was collected via vacuum filtration and subsequently washed with water and ether. The desired product (80% pure) was carried to the next step without further purification (MS (ESI-LCMS) for $C_{22}H_{19}N_3O_3$: 375.2 (MH⁺). To a solution of this material in THF:H₂O (3:1), was added lithium hydroxide (13.85 g, 330 mmol). The reaction was stirred at room temperature and monitored by LCMS. The reaction was complete after 3 h. The reaction mixture was poured into 1N HCl (500 mL) and shaken vigorously. Vacuum filtration was used to collect the resulting yellow solid, which was confirmed by LCMS. Extraction of the aqueous layer with ethyl acetate (3x, 1L) was used to recover the remaining product. The combined organic layers were washed with brine (3x, 200 mL), dried over anhydrous sodium sulfate, filtered and concentrated in vacuo to afford a yellow solid (MS (ESI-LCMS) for $C_{21}H_{17}N_3O_3$: 360.2 (MH⁺). To a portion of this yellow solid (2.0 g, 5.4 mmol) in DMF (50 mL) was added HATU (3.1 g, 8.1 mmol), *N,N*-diisopropylethylamine (2.8 mL, 16 mmol) and (*S*)-3-amino-1-*N*-Boc-pyrrolidine (1.5 g, 8.1 mmol). The reaction mixture was stirred at room temperature and monitored by LCMS. The reaction was complete after 1 h. The reaction mixture was diluted with water (200 mL), extracted with ethyl acetate (3x, 300 mL), washed with 5 % LiCl (3x, 100 mL) and brine (3x, 200 mL), dried over anhydrous sodium sulfate, filtered and concentrated under reduced pressure.

The material was purified by silica gel chromatography to afford a yellow solid (2.2 g, 92 % yield) (MS (ESI-LCMS) for $C_{30}H_{34}N_6O_{34}$: 443.2 (MH⁺). This material was dissolved in methanol (10 mL), and 4M HCl in dioxane (10 mL) was added. The reaction mixture was stirred at room temperature and monitored by LCMS. The reaction was complete after 2 h. The reaction mixture was concentrated in vacuo to afford a yellow solid as an HCl salt. ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.1 (d, 1H), 8.95 (s, 1H), 8.75 (d, 1H), 8.65 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.3-7.2 (m, 4H), 5.6 (q, 1H), 4.4 (m, 1H), 3.1-2.8 (m, 6H), 2.0 (m, 2H), 1.7 (m, 2H), 1.85 (m, 2H). MS (EI) for $C_{25}H_{26}N_6O_2$: 443 (MH⁺).

[0976] 3-Amino-6-{3-[(benzylamino)carbonyl]phenyl}-*N*-methoxy-*N*-methylpyrazine-2-carboxamide: 3-Carboxyphenylboronic acid (10 g, 61 mmol) was dissolved in thionyl chloride (60 mL) and DMF (catalytic), and the mixture was stirred at room temperature for 12 h. Thionyl chloride and DMF were removed under reduced pressure and the acid chloride was isolated as a white solid. A THF (100 mL) solution of the acid chloride was added dropwise to a MeOH (100 mL) solution of benzylamine (7 mL, 64 mmol) and triethylamine (18 mL), and the mixture was stirred at room temperature for 2 h. The solvent was removed under reduced pressure to give the benzamide as a white solid (MS (EI) for $C_{14}H_{14}BN_2O_3$: 256 (MH⁺). A mixture containing this benzamide (15 g, 59 mmol), methyl-3-amino-6-bromopyrazine carboxylate (15 g, 66 mmol), tris(dibenzylideneactone)dipalladium (1.4g, 1.5mmol), 1,1'-bis(diphenylphosphine) ferrocene (1.0 g, 1.9 mmol), and triethylamine (100 mL, 720 mmol) in DMF (200 mL) and water (30 mL), was heated at 80 °C for 48 h. The reaction mixture was cooled to room temperature and diluted with water and ethyl acetate. The layers were separated and the organic layer was extracted 2x with 5% aqueous lithium chloride, dried over sodium sulfate, filtered, and concentrated under reduced pressure to give a brown solid. The product was purified by triturating the brown solid with dichloromethane (100 mL) and filtering the resultant mixture to give the methyl ester as a yellow solid (3.0 g, 15% yield). *N,O*-Dimethylhydroxylamine was dissolved in anhydrous dichloromethane (15 mL). Trimethylaluminum (7 mL of a 2.0 M solution in hexanes) was added dropwise at 0 °C under N₂ and the solution was warmed to room temperature and stirred for 30 min. The previously prepared methyl ester (1.0 g, 2.8 mmol) was stirred in anhydrous dichloromethane (40 mL) and this solution was added in small portions to the reaction under N₂. The resulting solution was stirred at room temperature for 12 h. The reaction

mixture was acidified with 0.5 N aqueous hydrochloric acid and extracted 3x with dichloromethane. The combined organic layers were washed with brine, dried over sodium sulfate, filtered, and concentrated under reduced pressure to give the product as a yellow solid (0.82 g, 76% yield). ^1H NMR (400 MHz, d_6 -DMSO): δ 9.23 (t, 1H), 8.50 (s, 1H), 8.16 (d, 1H), 7.93 (d, 1H), 7.62 (t, 1H), 7.43-7.38 (m, 5H), 7.35-7.30 (m, 1H), 6.82 (br s, 2H), 4.58 (d, 2H), 3.76 (s, 3H), 3.40 (s, 3H). MS (EI) for $\text{C}_{20}\text{H}_{21}\text{N}_5\text{O}_3$: 392 (MH^+).

[0977] 3-{5-Amino-6-[(3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-

(phenylmethyl)benzamide: 3-Fluorophenylmagnesium bromide (1.5 mL of a 0.5 M solution in THF, 0.75 mmol) was added dropwise to a THF (10 mL) solution of 3-amino-6-{3-[(benzylamino)carbonyl]phenyl}-N-methoxy-N-methylpyrazine-2-carboxamide (0.10 g, 0.25 mmol) at 0 °C under N_2 . The reaction mixture was warmed to room temperature and stirred for 2 h. The mixture was diluted with water and extracted 3x with ethyl acetate. The combined organic layers were dried with magnesium sulfate, filtered, and concentrated under reduced pressure to give a yellow oil. The crude product was separated by preparative HPLC and lyophilized to give a yellow solid (4 mg, 4% yield). ^1H NMR (400 MHz, d_6 -DMSO): δ 9.10 (t, 1H), 9.05 (s, 1H), 8.42 (bs, 1H), 8.01 (m, 2H), 7.86-7.80 (m, 2H), 7.74 (d, 1H), 7.57-7.51 (m, 2H), 7.47-7.43 (m, 2H), 7.38-7.32 (m, 2H), 7.28-7.24 (m, 2H), 6.47-6.59 (m, 1H), 4.50 (d, 2H). MS (EI) for $\text{C}_{25}\text{H}_{19}\text{FN}_4\text{O}_2$: 427 (MH^+).

[0978] 3-(5-Amino-6-[(1E)-N-[4-(methyloxy)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide: A solution of 3-amino-6-{3-[(benzylamino)carbonyl]phenyl}-N-methoxy-N-methylpyrazine-2-carboxamide (0.10 g, 0.25 mmol) in THF (5 mL) was cooled to 0 °C, and a solution of CH_3MgCl (0.75 mL, 1 M in THF, 3.0 equiv.) was added slowly. The mixture was allowed to warm to room temperature over 1.5 hr. The mixture was then cooled to 0 °C and quenched slowly with saturated NH_4Cl solution. The mixture was extracted with ethyl acetate, washed with brine then water. The ethyl acetate extract was evaporated to afford the corresponding methyl ketone as a yellow solid. To a portion of this residue (35 mg, 0.1 mmol) in ethanol in a pressure tube (5 mL) was added 4-methoxyphenylhydrazine hydrochloride (52 mg, 3 mmol). This mixture was then heated to for 12 h. The solvent was removed under reduced pressure and the crude mixture was dissolved in a minimal amount of methanol and run through a plug of silica gel to give the product as a white solid (33 mg,

70 % yield). ^1H NMR (400 MHz, CDCl_3): δ 8.40 (m, 1H), 8.0 (m, 2H), 7.80 (m, 2H), 7.25 (m, 1H), 7.2 (m, 4H), 6.9 (m, 4H), 4.8 (s, 2H), 3.8 (s, 3H), 2.6 (s, 3H). MS (EI) for $\text{C}_{27}\text{H}_{26}\text{N}_6\text{O}_2$: 467 (MH^+).

[0979] 3-(5-Amino-6-formylpyrazin-2-yl)-*N*-benzylbenzamide: A solution of 3-amino-6-{3-[(benzylamino)carbonyl]phenyl}-*N*-methoxy-*N*-methylpyrazine-2-carboxamide (2.5 g, 6.4 mmol) in THF (20 mL) was cooled to 0 °C. Under N_2 lithium aluminum hydride (0.364g, 9.58 mmol) was added and the reaction was stirred at 0 °C for 1 h. The mixture was then quenched with H_2O at 0 °C. Removal of solids via filtration, concentration of the filtrate and dissolution of the solid residue in H_2O followed by washing 3x with dichloromethane gave the product (1.90 g, 89 % yield) with sufficient purity to be used without further purification.

[0980] 3-[5-Amino-6-(morpholin-4-ylmethyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 3-(5-Amino-6-formylpyrazin-2-yl)-*N*-benzylbenzamide (1.27 g, 3.83 mmol) was dissolved in dichloroethane (30 mL) at 0 °C. Morpholine (400 mg, 4.6 mmol), and $\text{NaBH}(\text{OAc})_3$ (2.03 g, 9.58 mmol) were added and the reaction was stirred at room temperature for 20 h. The reaction mixture was diluted with saturated sodium carbonate and extracted 3x with ethyl acetate. The combined ethyl acetate layers were dried over sodium sulfate, filtered, and concentrated *in vacuo* to afford the product (1.41 g, 88 % yield). ^1H NMR (400 MHz, d_4 - CD_3OD): δ 8.60 (s, 1H), 8.47 (t, 1H), 8.12 (m, 1H), 7.84 (m, 1H), 7.55 (t, 1H), 7.31-7.89 (m, 4H), 7.23-7.26 (m, 1H), 4.61 (s, 2H), 4.43 (s, 2H), 3.32-4.16 (m, 8H). MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_5\text{O}_2$: 404 (MH^+).

[0981] Using the same or similar synthetic techniques, substituting with the appropriate reagents such as the respective amines, the following compounds of the invention were prepared:

[0982] 3-{5-Amino-6-[(3,5-difluorophenyl)carbonyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.25 (m, 2H), 8.45 (s, 1H), 8.08 (s, 1H), 8.05 (m, 1H), 8.02 (m, 1H), 7.67-7.53 (m, 5H), 7.35-7.24 (M, 5H), 4.51 (M, 2H). MS (EI) for $\text{C}_{25}\text{H}_{18}\text{F}_2\text{N}_4\text{O}_2$: 445 (MH^+).

[0983] 3-[5-Amino-6-(biphenyl-4-ylcarbonyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.06 (t, 1H), 8.99 (s, 1H), 8.42 (s, 1H), 8.06-8.01 (m, 3H), 7.91 (s, 2H), 7.81-7.78 (m, 3H), 7.73-7.71 (m, 2H), 7.52-7.44 (m, 3H), 7.40-7.36 (m,

1H), 7.27-7.20 (m, 4H), 7.16-7.12 (m, 1H), 4.35 (d, 2H). MS (EI) for $C_{31}H_{24}N_4O_2$: 485 (MH^+).

[0984] 3-{5-Amino-6-[(4-chloro-3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.25 (m, 2H), 8.48 (s, 1H), 8.09 (s, 1H), 8.05 (m, 1H), 8.02 (m, 1H), 7.71-7.51 (m, 5H), 7.35-7.24 (m, 5H), 4.51 (m, 2H). MS (EI) for $C_{25}H_{18}ClFN_4O_2$: 461 (MH^+).

[0985] 3-(5-Amino-6-{[2,4-bis(methyloxy)phenyl]carbonyl}pyrazin-2-yl)-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.11 (m, 1H), 8.95 (s, 1H), 8.36 (s, 1H), 7.91-7.83 (m, 4H), 7.53-7.23 (m, 7H), 6.68-6.62 (m, 2H), 4.50 (m, 2H), 3.85 (s, 3H), 3.65 (s, 3H). MS (EI) for $C_{27}H_{24}N_4O_4$: 469 (MH^+).

[0986] 3-(5-Amino-6-{[4-(dimethylamino)phenyl]carbonyl}pyrazin-2-yl)-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.11 (t, 1H), 8.90 (s, 1H), 8.45 (s, 1H), 8.44 (d, 1H), 8.07 (d, 2H), 7.98 (d, 1H), 7.55 (t, 3H), 7.31-7.30 (m, 4H), 7.28-7.23 (m, 1H), 6.75 (d, 2H), 4.50 (d, 2H), 3.03 (s, 6H). MS (EI) for $C_{27}H_{25}N_5O_2$: 452 (MH^+).

[0987] 3-{5-Amino-6-[(4-methylphenyl)acetyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.18 (t, 1H), 9.02 (s, 1H), 8.59 (s, 1H), 8.26 (d, 1H), 7.91 (d, 2H), 7.61 (t, 1H), 7.38-7.29 (m, 4H), 7.24-7.17 (m, 3H), 7.08-7.06 (m, 3H), 4.53-4.51 (m, 4H), 2.23 (s, 3H). MS (EI) for $C_{27}H_{24}N_4O_2$: 437 (MH^+).

[0988] 1,1-Dimethylethyl 4-([3-amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)piperidine-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 9.16 (t, 1H), 8.9 (s, 1H), 8.55 (d, 1H), 8.5 (s, 1H), 8.3 (d, 1H), 7.82 (d, 1H), 7.8-7.6 (m, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 4.0 (m, 3H), 3.0-2.7 (m, 2H), 1.8 (d, 2H), 1.6 (q, 2H), 1.4 (s, 9H). MS(EI) for $C_{29}H_{34}N_6O_4$: 431 (MH^+).

[0989] 1,1-Dimethylethyl 4-([3-amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)methylpiperidine-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 9.17 (t, 1H), 8.92 (t, 1H), 8.9 (s, 1H), 8.5 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (d, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 3.9 (m, 2H), 3.2 (t, 2H), 2.65 (m, 2H), 1.8-1.6 (m, 3H), 1.4 (s, 9H), 1.05 (m, 2H). MS(EI) for $C_{30}H_{36}N_6O_4$: 445 (MH^+).

- [0990] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-piperidin-4-ylpyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (t, 1H), 8.9 (s, 1H), 8.8 (d, 1H), 8.65 (m, 1H), 8.3 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 4.05 (m, 1H), 3.35 (m, 2H), 3.0 (m, 2H), 2.0-1.8 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431 (MH^+).
- [0991] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-(piperidin-4-ylmethyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.42 (t, 1H), 9.15 (t, 1H), 8.9 (s, 1H), 8.62 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.4-7.2 (m, 5H), 4.5 (d, 2H), 3.3-3.2 (m, 4H), 2.9-2.7 (m, 2H), 1.95-1.75 (m, 3H), 1.5-1.35 (m, 2H). MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2$: 445 (MH^+).
- [0992] 3-[5-Amino-6-(morpholin-4-ylcarbonyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.18 (t, 1H), 8.78 (s, 1H), 8.42 (s, 1H), 8.1 (d, 1H), 7.88 (d, 1H), 7.55 (m, 1H), 7.38-7.24 (m, 5H), 4.55 (d, 2H), 3.78-3.4 (m, 8H). MS (EI) for $\text{C}_{23}\text{H}_{23}\text{N}_5\text{O}_3$: 418 (MH^+).
- [0993] 3-(5-Aminopyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.6 (s, 1H), 8.45 (s, 1H), 8.1 (d, 1H), 8.0 (s, 1H), 7.85 (d, 1H), 7.55 (t, 1H), 7.4-7.3 (m, 5H), 6.65 (s, 2H). MS (EI) for $\text{C}_{18}\text{H}_{16}\text{N}_4\text{O}$: 305 (MH^+).
- [0994] 1,1-Dimethylethyl 3-([(3-amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.18 (t, 1H), 8.95 (s, 1H), 8.75 (m, 1H), 8.55 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.4-7.3 (m, 4H), 7.25 (m, 1H), 4.55 (m, 3H), 3.6 (m, 1H), 3.45 (m, 1H), 3.3 (m, 2H), 2.15-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{28}\text{H}_{32}\text{N}_6\text{O}_4$: 417 (MH^+).
- [0995] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.55 (m, 2H), 9.3 (s, 1H), 8.95 (s, 1H), 8.75 (s, 1H), 8.35 (d, 1H), 7.95 (d, 1H), 7.55 (t, 1H), 7.4-7.3 (m, 4H), 7.25 (t, 1H), 4.7 (m, 1H), 4.5 (d, 2H), 3.4-3.2 (m, 2H), 2.3-2.1 (m, 2H), 1.1 (t, 2H). MS (EI) for $\text{C}_{23}\text{H}_{24}\text{N}_6\text{O}_2$: 417 (MH^+).
- [0996] 3-[5-Amino-6-[(4-methylpiperazin-1-yl)carbonyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.75 (s, 1H), 8.45 (s, 1H), 8.18 (d, 1H), 7.85 (d, 1H), 7.55 (t, 1H), 7.35-7.2 (m, 5H), 6.72 (s, 2H), 4.5 (d, 2H), 3.7 (t, 2H), 3.5 (t, 2H), 2.45 (t, 2H), 2.35 (t, 2H), 2.2 (s, 3H). MS (EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431 (MH^+).

- [0997] 3-(5-Amino-6-{[4-(4-fluorophenyl)piperazin-1-yl]carbonyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.75 (s, 1H), 8.45 (t, 1H), 8.18 (tt, 1H), 7.85 (tt, 1H), 5.75 (t, 1H), 7.3-7.2 (m, 5H), 7.0 (m, 4H), 6.8 (s, 2H), 4.5 (d, 2H), 3.82 (t, 2H), 3.65 (t, 2H), 3.25 (t, 2H), 3.15 (t, 2H). MS (EI) for $\text{C}_{29}\text{H}_{27}\text{N}_6\text{O}_2\text{F}$: 511 (MH^+).
- [0998] 3-(5-Amino-6-{[4-(phenylmethyl)piperazin-1-yl]carbonyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.7 (s, 1H), 8.4 (t, 1H), 8.05 (d, 1H), 7.85 (d, 1H), 7.55 (t, 1H), 7.35-7.2 (m, 10H), 6.7 (s, 2H), 4.5 (d, 2H), 3.7 (t, 2H), 3.5 (m, 4H), 2.5-2.4 (m, 4H). MS (EI) for $\text{C}_{30}\text{H}_{30}\text{N}_6\text{O}_2$: 507 (MH^+).
- [0999] Methyl *N*-{[3-amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl}glycinate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.22 (t, 1H), 9.15 (t, 1H), 8.92 (s, 1H), 8.5 (s, 1H), 8.36 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 4.05 (d, 2H), 3.65 (s, 3H). MS (EI) for $\text{C}_{22}\text{H}_{21}\text{N}_5\text{O}_4$: 420 (MH^+).
- [1000] *N*-{[3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl}glycine: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.93 (s, 1H), 8.7 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.55 (d, 2H), 4.0 (d, 2H). MS (EI) for $\text{C}_{21}\text{H}_{19}\text{N}_5\text{O}_4$: 406 (MH^+).
- [1001] Methyl 1-{[3-amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl}-*L*-prolinate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (m, 1H), 8.85-8.75 (s, 1H), 8.45-8.35 (s, 1H), 8.12-7.95 (d, 1H), 7.88 (d, 1H), 7.6 (m, 1H), 7.4-7.25 (m, 5H), 5.25 (m, 1H), 4.5 (m, 4H), 4.0 (t, 1H), 3.6 (s, 3H), 2.4-2.25 (m, 1H), 2.0-1.75 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{25}\text{N}_5\text{O}_4$: 460 (MH^+).
- [1002] 3-[(6a*S*)-6,11-dioxo-5,6a,7,8,9,11-hexahydro-6*H*-pyrazino[2,3-*e*]pyrrolo[1,2-*a*][1,4]diazepin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.3 (s, 1H), 9.3 (s, 1H), 9.2 (t, 1H), 8.7 (s, 1H), 8.3 (d, 1H), 8.0 (d, 1H), 7.7 (t, 1H), 6.4-6.2 (m, 5H), 4.5 (m, 3H), 3.7 (m, 1H), 3.5 (m, 1H), 3.0 (s, 1H), 2.0-1.8 (m, 3H). MS (EI) for $\text{C}_{24}\text{H}_{21}\text{N}_5\text{O}_3$: 428 (MH^+).
- [1003] Methyl 1-{[3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl}piperidine-3-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.75 (d, 1H), 8.42 (d, 1H), 8.08 (t, 1H), 7.85 (d, 1H), 7.55 (m, 1H), 7.35-7.25 (m, 5H), 6.72 (d, 2H), 4.5 (d, 2H), 4.1-3.9 (m, 1H), 3.65 (m, 2H), 3.45 (s, 2H), 3.25-3.05 (m, 2H), 2.75 (m, 1H), 2.0 (m, 1H), 1.8-1.5 (m, 3H). MS (EI) for $\text{C}_{26}\text{H}_{27}\text{N}_5\text{O}_4$: 474 (MH^+).

- [1004] 1,1-Dimethylethyl 4-{{[3-amino-6-(3-{{[(phenylmethyl)amino]carbonyl}phenyl}pyrazin-2-yl]carbonyl}piperazine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.72 (s, 1H), 8.6 (s, 1H), 8.05 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 6.75 (s, 2H), 4.5 (d, 2H), 3.65 (t, 2H), 3.5-3.35 (m, 6H), 1.4 (s, 9H). MS(EI) for $\text{C}_{28}\text{H}_{32}\text{N}_6\text{O}_4$: 417 (MH^+).
- [1005] 3-[5-Amino-6-(piperazin-1-ylcarbonyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.75 (br s, 2H), 9.1 (t, 1H), 8.8 (s, 1H), 8.45 (s, 1H), 8.05 (d, 1H), 7.86 (d, 1H), 7.55 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 3.95 (t, 2H), 3.8 (t, 2H), 3.25 (t, 2H), 3.15 (t, 2H). MS (EI) for $\text{C}_{23}\text{H}_{24}\text{N}_6\text{O}_2$: 417 (MH^+).
- [1006] 3-(5-Amino-6-cyanopyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.16 (m, 1H), 8.99 (s, 1H), 8.46-8.43 (m, 1H), 8.13-8.09 (m, 1H), 7.94-7.90 (m, 1H), 7.61-7.52 (m, 3H), 7.37-7.31 (m, 4H), 7.28-7.22 (m, 1H), 4.55-4.49 (m, 2H). MS (EI) for $\text{C}_{19}\text{H}_{15}\text{N}_5\text{O}$: 330 (MH^+).
- [1007] *N*-{{[3-Amino-6-(3-{{[(phenylmethyl)amino]carbonyl}phenyl}pyrazin-2-yl]carbonyl}-L-glutamic acid: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.95 (s, 1H), 8.9 (d, 1H), 8.55 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 4.45 (t, 1H), 2.35 (m, 2H), 2.3-2.05 (m, 2H). MS (EI) for $\text{C}_{24}\text{H}_{23}\text{N}_5\text{O}_6$: 478 (MH^+).
- [1008] Ethyl *N*-{{[3-amino-6-(3-{{[(phenylmethyl)amino]carbonyl}phenyl}pyrazin-2-yl]carbonyl}-L-tyrosinate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.98 (s, 1H), 8.68 (d, 1H), 8.58 (s, 1H), 8.22 (d, 1H), 7.92 (d, 1H), 7.65 (t, 1H), 7.4-7.2 (m, 5H), 7.12 (d, 2H), 6.65 (d, 2H), 4.6 (m, 4H), 4.15 (m, 4H), 3.1 (d, 2H), 1.2 (t, 3H). MS (EI) for $\text{C}_{30}\text{H}_{29}\text{N}_5\text{O}_5$: 540 (MH^+).
- [1009] *N*-{{[3-Amino-6-(3-{{[(phenylmethyl)amino]carbonyl}phenyl}pyrazin-2-yl]carbonyl}tyrosine: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.22 (t, 1H), 8.94 (s, 1H), 8.64 (d, 1H), 8.56 (s, 1H), 8.18 (d, 1H), 7.92 (d, 1H), 7.62 (t, 1H), 7.38-7.22 (m, 5H), 7.1 (d, 2H), 6.64 (d, 2H), 4.62-4.44 (m, 3H), 3.12 (d, 2H). MS (EI) for $\text{C}_{28}\text{H}_{25}\text{N}_5\text{O}_5$: 512 (MH^+).
- [1010] 1,1-Dimethylethyl [3-{{[3-amino-6-(3-{{[(phenylmethyl)amino]carbonyl}phenyl}pyrazin-2-yl]carbonyl}amino)propyl]carbamate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.16 (t, 1H), 9.0 (t, 1H), 8.93 (s, 1H), 8.54 (s, 1H), 8.44 (d, 1H), 7.9 (d, 1H), 7.58 (t, 1H), 7.38-7.23 (m, 5H), 6.9 (t, 1H), 4.55 (d, 2H), 3.35 (m, 2H), 3.0 (q, 2H), 1.65 (q, 2H), 1.38 (s, 9H). MS (EI) for $\text{C}_{27}\text{H}_{32}\text{N}_6\text{O}_4$: 405 (MH^+).

- [1011] 3-Amino-*N*-(3-aminopropyl)-6-(3-([(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.55 (t, 1H), 9.3 (t, 1H), 8.92 (s, 1H), 8.7 (s, 1H), 8.35 (m, 1H), 8.0 (s, 2H), 7.9 (m, 1H), 7.55 (t, 1H), 7.4-7.2 (m, 5H), 4.5 (d, 2H), 3.45 (m, 2H), 2.85 (m, 2H), 1.85 (m, 2H). MS (EI) for $\text{C}_{22}\text{H}_{24}\text{N}_6\text{O}_2$: 405 (MH^+).
- [1012] 1-([3-Amino-6-(3-([(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl)piperidine-3-carboxylic acid: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.7 (s, 1H), 8.45 (s, 1H), 8.05 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 6.7 (s, 2H), 4.5 (d, 2H), 4.45-4.2 (m, 1H), 3.9-3.6 (m, 1H), 3.3 (m, 1H), 3.1 (m, 1H), 2.6 (m, 1H), 2.0 (m, 1H), 1.8-1.5 (m, 3H). MS(EI) for $\text{C}_{25}\text{H}_{25}\text{N}_5\text{O}_4$: 460 (MH^+).
- [1013] 1,1-Dimethylethyl [2-([3-amino-6-(3-([(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl)amino)ethyl]carbamate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.95 (s, 1H), 8.95 (t, 1H), 8.55 (s, 1H), 8.42 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 7.02 (t, 1H), 4.5 (d, 2H), 3.4 (m, 2H), 3.2 (m, 2H), 1.4 (s, 9H). MS(EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}_4$: 391 (MH^+).
- [1014] 3-Amino-*N*-(2-aminoethyl)-6-(3-([(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.6 (t, 1H), 9.3 (t, 1H), 8.95 (s, 1H), 8.75 (s, 1H), 8.37 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 3.62 (t, 2H), 3.04 (t, 2H). MS(EI) for $\text{C}_{21}\text{H}_{22}\text{N}_6\text{O}_2$: 391 (MH^+).
- [1015] 1,1-Dimethylethyl (1-([3-amino-6-(3-([(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl)piperidin-4-yl)carbamate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.75 (s, 1H), 8.4 (s, 1H), 8.07 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 6.95 (d, 1H), 6.7 (s, 2H), 4.5 (d, 2H), 4.4 (m, 1H), 3.8 (m, 1H), 3.6 (m, 1H), 3.2 (m, 1H), 3.05 (m, 1H), 1.9-1.7 (m, 2H), 1.5-1.3 (m, 2H), 1.4 (s, 9H). MS(EI) for $\text{C}_{29}\text{H}_{34}\text{N}_6\text{O}_4$: 431 (MH^+).
- [1016] 3-{5-Amino-6-[(4-aminopiperidin-1-yl)carbonyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.23 (t, 1H), 8.75 (s, 1H), 8.42 (s, 1H), 8.25 (t, 3H), 8.05 (d, 1H), 7.95 (d, 1H), 7.57 (t, 1H), 7.4-7.2 (m, 5H), 4.5 (d, 2H), 3.9 (m, 1H), 3.7-3.5 (m, 1H), 3.35 (m, 1H), 3.2 (m, 1H), 2.95 (m, 1H), 2.1-1.9 (m, 2H), 1.7-1.5 (t, 2H). MS(EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431 (MH^+).
- [1017] 3-Amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl}phenyl)-*N*-[(3*R*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.1 (d, 1H),

8.95 (s, 1H), 8.75 (d, 1H), 8.55 (s, 1H), 8.35 (d, 1H), 7.93 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 4.4 (m, 1H), 3.1-3.0 (m, 3H), 2.9-2.8 (m, 3H), 2.15-2.0 (m, 2H), 1.8 (m, 2H). MS (EI) for $C_{25}H_{26}N_6O_2$: 443 (MH^+).

[1018] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-N-[(3S)-1-(furan-2-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.96 (s, 1H), 8.9 (d, 1H), 8.85 (q, 1H), 8.6 (s, 1H), 8.35 (d, 1H), 7.95-7.85 (q, 2H), 7.6 (m, 1H), 7.3-7.15 (m, 4H), 6.65 (m, 1H), 5.6 (q, 1H), 4.65-4.5 (m, 1H), 4.15-3.95 (m, 1H), 3.8 (m, 2H), 3.6 (m, 1H), 3.05-2.8 (m, 2H), 2.3-2.0 (m, 4H). MS (EI) for $C_{30}H_{28}N_6O_4$: 537 (MH^+).

[1019] N-[3-(Acetylamino)propyl]-3-amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 9.0 (t, 1H), 8.95 (s, 1H), 8.6 (s, 1H), 8.4 (d, 1H), 7.95 (t, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.4-7.25 (m, 5H), 4.55 (d, 2H), 3.35 (q, 2H), 3.15 (q, 2H), 1.8 (s, 3H), 1.65 (m, 2H). MS (EI) for $C_{24}H_{26}N_6O_3$: 447 (MH^+).

[1020] 3-Amino-N-{3-[(furan-2-ylcarbonyl)amino]propyl}-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.16 (t, 1H), 9.05 (t, 1H), 8.95 (s, 1H), 8.6 (s, 1H), 8.5 (t, 1H), 8.44 (d, 1H), 7.92 (d, 1H), 7.84 (s, 1H), 7.6 (t, 1H), 7.35-7.24 (m, 5H), 7.1 (d, 1H), 6.62 (m, 1H), 4.54 (d, 2H), 3.42-3.28 (m, 4H), 1.78 (m, 2H). MS (EI) for $C_{27}H_{26}N_6O_4$: 499 (MH^+).

[1021] 3-Amino-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-N-[(3S)-1-(pyridin-4-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.95-8.9 (m, 2H), 8.85-8.75 (dd, 1H), 8.7 (m, 2H), 8.55 (d, 1H), 8.45 (s, 1H), 7.95 (t, 1H), 7.6 (t, 1H), 7.5 (d, 1H), 7.45 (d, 1H), 7.25 (m, 4H), 5.6 (q, 1H), 4.55-4.5 (m, 1H), 3.9-3.4 (m, 3H), 3.1-2.9 (m, 3H), 2.25-1.95 (m, 3H), 1.35 (m, 2H). MS (EI) for $C_{31}H_{29}N_7O_3$: 548 (MH^+).

[1022] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-N-{3-[(pyridin-4-ylcarbonyl)amino]propyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 9.05 (t, 1H), 8.95 (s, 1H), 8.88 (t, 1H), 8.7 (d, 2H), 8.58 (s, 1H), 8.45 (d, 1H), 7.92 (d, 1H), 7.75 (d, 2H), 7.6 (t, 1H), 7.35-7.25 (m, 5H), 4.55 (d, 2H), 3.4 (m, 4H), 1.85 (m, 2H). MS (EI) for $C_{28}H_{27}N_7O_3$: 510 (MH^+).

[1023] N-[3-([3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino]propyl]quinoxaline-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO):

δ 9.45 (s, 1H), 9.2 (t, 1H), 9.1 (t, 1H), 9.05 (t, 1H), 8.9 (s, 1H), 8.5 (s, 1H), 8.4 (d, 1H), 8.15 (m, 2H), 7.95 (m, 2H), 7.85 (d, 1H), 7.55 (t, 1H), 7.35-7.15 (m, 5H), 4.5 (d, 2H), 3.5-3.4 (m, 4H), 1.85 (m, 2H). MS (EI) for $C_{31}H_{28}N_8O_3$: 561 (MH^+).

[1024] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-[3-([(6-(trifluoromethyl)pyridin-3-yl)carbonyl]amino)propyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 2H), 9.0 (m, 2H), 8.9 (s, 1H), 8.55 (s, 1H), 8.4 (m, 2H), 7.95 (d, 1H), 7.35 (d, 1H), 7.55 (t, 1H), 7.35-7.15 (m, 5H), 4.5 (d, 2H), 3.4 (m, 4H), 1.8 (m, 2H). MS (EI) for $C_{29}H_{26}N_7O_3F_3$: 578 (MH^+).

[1025] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-{3-[(quinolin-8-ylsulfonyl)amino]propyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 9.05 (m, 1H), 8.9 (s, 1H), 8.8 (t, 1H), 8.5 (m, 2H), 8.35 (q, 2H), 8.25 (d, 1H), 7.9 (d, 1H), 7.7 (m, 2H), 7.55 (t, 1H), 7.35-7.2 (m, 5H), 4.55 (d, 2H), 3.25 (q, 2H), 2.85 (q, 2H), 1.6 (q, 2H). MS (EI) for $C_{31}H_{29}N_7O_4S$: 596 (MH^+).

[1026] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-{3-[(pyridin-2-ylsulfonyl)amino]propyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (t, 1H), 8.95 (m, 2H), 8.7 (d, 1H), 8.6 (s, 1H), 8.35 (d, 1H), 8.05 (t, 1H), 7.9 (m, 3H), 7.65-7.55 (m, 2H), 7.4-7.25 (m, 5H), 4.55 (d, 2H), 3.3 (q, 2H), 3.0 (q, 2H), 1.7 (q, 2H). MS (EI) for $C_{27}H_{27}N_7O_4S$: 546 (MH^+).

[1027] 3-Amino-*N*-[3-([(3,5-dimethylisoxazol-4-yl)amino]carbonyl)amino]propyl]-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 9.05 (t, 1H), 8.95 (s, 1H), 8.55 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.6 (s, 1H), 7.5 (t, 1H), 7.35-7.2 (m, 5H), 6.35 (t, 1H), 4.55 (d, 2H), 3.35 (m, 2H), 3.15 (q, 2H), 2.25 (s, 3H), 2.05 (s, 3H), 1.65 (q, 2H). MS (EI) for $C_{28}H_{30}N_8O_4$: 543 (MH^+).

[1028] 3-Amino-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)-*N*-{2-[(pyridin-4-ylcarbonyl)amino]ethyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 9.05 (t, 1H), 9.0 (t, 1H), 8.95 (s, 1H), 8.7 (d, 2H), 8.6 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.75 (d, 2H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 3.5 (t, 4H). MS(EI) for $C_{27}H_{29}N_7O_3$: 496 (MH^+).

[1029] 3-Amino-*N*-{2-[(furan-2-ylcarbonyl)amino]ethyl}-6-(3-([(phenylmethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 9.05 (t, 1H), 8.95 (s, 1H), 8.65 (t, 1H), 8.6 (s, 1H), 8.4 (d, 1H),

7.9 (d, 1H), 7.82 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 7.06 (d, 1H), 6.6 (d, 1H), 4.5 (d, 2H), 3.5 (t, 4H). MS(EI) for $C_{26}H_{24}N_6O_4$: 485 (MH^+).

[1030] *N,N'*-Cyclohexane-1,2-diylbis[3-amino-6-(3-

{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 2H), 8.9 (s, 2H), 8.75 (d, 2H), 8.57 (s, 2H), 8.4 (d, 2H), 7.9 (d, 2H), 7.57 (t, 2H), 7.35-7.2 (m, 10H), 4.55 (d, 4H), 4.0 (m, 2H), 2.05 (d, 2H), 1.8 (d, 2H), 1.6 (m, 2H), 1.4 (t, 2H). MS(EI) for $C_{44}H_{42}N_{10}O_4$: 775 (MH^+).

[1031] *N*-[2-(Acetylamino)ethyl]-3-amino-6-(3-

{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.25 (t, 1H), 8.98 (t, 1H), 8.95 (s, 1H), 8.6 (s, 1H), 8.4 (d, 1H), 8.16 (t, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.55 (d, 2H), 3.4 (t, 2H), 3.27 (t, 2H), 1.8 (s, 3H). MS(EI) for $C_{23}H_{24}N_6O_3$: 433 (MH^+).

[1032] 3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)-*N*-{2-[(quinolin-8-

ylsulfonyl)amino]ethyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.9 (s, 1H), 8.8 (d, 1H), 8.7 (t, 1H), 8.5 (s, 1H), 8.3 (m, 3H), 8.2 (d, 1H), 7.9 (d, 1H), 7.65 (t, 1H), 7.57 (t, 1H), 7.4-7.2 (m, 7H), 4.55 (d, 2H), 3.35 (t, 2H), 3.05 (t, 2H). MS(EI) for $C_{30}H_{27}N_7O_4S$: 582 (MH^+).

[1033] *N*-[2-({[3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-

yl]carbonyl}amino)ethyl]quinoxaline-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.45 (s, 1H), 9.35 (t, 1H), 9.15 (t, 1H), 9.07 (t, 1H), 8.95 (s, 1H), 8.57 (s, 1H), 8.4 (d, 1H), 8.2 (t, 2H), 8.0 (t, 2H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 3.6 (t, 4H). MS(EI) for $C_{30}H_{26}N_8O_3$: 547 (MH^+).

[1034] 3-Amino-*N*-(2-{[(2-chloropyridin-3-yl)carbonyl]amino}ethyl)-6-(3-

{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 9.0 (t, 1H), 8.95 (s, 1H), 8.8 (t, 1H), 8.6 (s, 1H), 8.45 (dd, 1H), 8.4 (d, 1H), 7.9 (m, 2H), 7.57 (t, 1H), 7.45 (m, 1H), 7.35-7.2 (m, 5H), 4.5 (d, 2H), 3.5 (t, 4H). MS(EI) for $C_{27}H_{24}N_7O_3Cl$: 530 (MH^+).

[1035] 3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)-*N*-{2-[(pyridin-2-

ylsulfonyl)amino]ethyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 8.9 (s, 1H), 8.85 (t, 1H), 8.63 (d, 1H), 8.5 (s, 1H), 8.37 (d, 1H), 8.1 (t, 1H), 8.0 (m, 2H), 7.9 (m, 2H), 7.57 (m, 2H), 7.35-7.2 (m, 5H), 4.55 (d, 2H), 3.4 (t, 2H), 3.2 (t, 2H). MS(EI) for $C_{26}H_{25}N_7O_4S$: 532 (MH^+).

- [1036] 3-Amino-*N*-[2-({[(3,5-dimethylisoxazol-4-yl)amino]carbonyl}amino)ethyl]-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 9.05 (t, 1H), 8.95 (s, 1H), 8.6 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.6 (s, 1H), 7.5 (t, 1H), 7.35-7.2 (m, 5H), 6.55 (t, 1H), 4.5 (d, 2H), 3.4 (t, 2H), 3.3 (t, 2H), 2.15 (s, 3H), 1.96 (s, 3H). MS(EI) for $\text{C}_{27}\text{H}_{28}\text{N}_8\text{O}_4$: 529 (MH^+).
- [1037] 3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.16 (m, 1H), 8.92 (s, 1H), 8.58-8.55 (m, 1H), 8.38-8.32 (m, 2H), 8.15-7.50 (bs, 2H), 7.79 (s, 1H), 7.60-7.54 (m, 1H), 7.38-7.31 (m, 4H), 7.28-7.22 (m, 1H), 4.58-4.53 (m, 2H). MS (EI) for $\text{C}_{19}\text{H}_{17}\text{N}_5\text{O}_2$: 348 (MH^+).
- [1038] 3-Amino-*N*-[3-({[4-(dimethylamino)phenyl]carbonyl}amino)propyl]-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (q, 2H), 8.95 (s, 1H), 8.6 (s, 1H), 8.45 (d, 1H), 8.25 (t, 1H), 7.9 (d, 1H), 7.7 (d, 2H), 7.6 (t, 1H), 7.35-7.2 (m, 5H), 6.65 (d, 2H), 4.45 (d, 2H), 3.35 (m, 4H), 2.95 (s, 6H), 1.75 (m, 2H). MS (EI) for $\text{C}_{31}\text{H}_{33}\text{N}_7\text{O}_3$: 552 (MH^+).
- [1039] 1,1-Dimethylethyl 4-[(3*S*)-3-({[3-amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl}amino)pyrrolidin-1-yl]piperidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.95 (s, 1H), 8.65 (d, 1H), 8.55 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.7 (s, 1H), 7.55 (t, 1H), 7.35-7.2 (m, 5H), 4.55 (d, 2H), 4.5 (m, 1H), 3.8 (d, 2H), 2.9-2.7 (m, 3H), 2.55 (m, 1H), 2.25-2.1 (m, 2H), 1.9-1.75 (m, 3H), 1.4 (s, 9H), 1.25 (m, 2H). MS (EI) for $\text{C}_{33}\text{H}_{41}\text{N}_7\text{O}_4$: 600 (MH^+).
- [1040] 3-Amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)-*N*-[(3*S*)-1-piperidin-4-ylpyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.65-9.45 (t, 1H), 9.3 (m, 2H), 8.98 (d, 1H), 8.75 (m, 2H), 8.5-8.3 (dd, 1H), 7.9 (t, 1H), 7.55 (m, 1H), 7.4-7.25 (m, 4H), 4.95-4.7 (m, 1H), 4.55 (m, 2H), 3.4 (m, 6H), 3.1 (m, 1H), 2.9 (m, 2H), 2.35-1.9 (m, 6H). MS (EI) for $\text{C}_{28}\text{H}_{33}\text{N}_7\text{O}_2$: 500 (MH^+).
- [1041] 1,1-Dimethylethyl (2*R*)-2-({[(3*S*)-3-({[3-amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]carbonyl}amino)pyrrolidin-1-yl]methyl}pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.93 (s, 1H), 8.6 (d, 1H), 8.55 (s, 1H), 8.25 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.55 (d, 2H), 4.4 (m, 1H), 3.8 (m, 1H), 3.2 (t, 2H), 2.8 (t, 2H), 2.6-2.4 (m, 4H), 2.2 (m, 1H), 1.9-1.7 (m, 5H), 1.4 (s, 9H). MS(EI) for $\text{C}_{33}\text{H}_{41}\text{N}_7\text{O}_4$: 600 (MH^+).

- [1042] 3-Amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[(3S)-1-[(2R)-pyrrolidin-2-ylmethyl]pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.4 (m, 1H), 9.7-9.4 (m, 3H), 9.35 (d, 1H), 8.96 (s, 1H), 8.8-8.7 (m, 1H), 8.5-8.3 (m, 1H), 7.9 (d, 1H); 7.57 (t, 1H), 7.35-7.2 (m, 5H), 5.0-4.7 (m, 1H), 4.5 (d, 2H), 4.0-3.6 (m, 3H), 3.2 (m, 3H), 2.6 (m, 1H), 2.4-2.1 (m, 3H), 2.0-1.6 (m, 3H). MS(EI) for $\text{C}_{28}\text{H}_{33}\text{N}_7\text{O}_2$: 500 (MH^+).
- [1043] 3-Amino-N-[(3S)-1-[[4-(dimethylamino)phenyl]methyl]pyrrolidin-3-yl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.9 (s, 1H), 8.6 (d, 1H), 8.57 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.35-7.2 (m, 5H), 7.1 (d, 2H), 6.63 (d, 2H), 4.55 (d, 2H), 4.4 (m, 1H), 3.45 (m, 2H), 2.82 (s, 6H), 2.7 (t, 2H), 2.4 (m, 1H), 2.2 (m, 1H), 1.82 (m, 1H). MS(EI) for $\text{C}_{32}\text{H}_{35}\text{N}_7\text{O}_2$: 550 (MH^+).
- [1044] 3-Amino-N-{2-[[[4-(dimethylamino)phenyl]amino]carbonyl]amino]ethyl}-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.15 (t, 1H), 9.05 (t, 1H), 8.95 (s, 1H), 8.6 (s, 1H), 8.42 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.45 (m, 2H), 7.35-7.2 (m, 5H), 6.57 (t, 1H), 4.5 (d, 2H), 3.4 (m, 4H), 3.05 (s, 6H). MS(EI) for $\text{C}_{30}\text{H}_{32}\text{N}_8\text{O}_3$: 553 (MH^+).
- [1045] 3-Amino-N-[2-[[[4-(dimethylamino)phenyl]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (t, 1H), 9.15 (t, 1H), 8.95 (s, 1H), 8.68 (s, 1H), 8.45 (m, 2H), 7.9 (d, 1H), 7.7 (d, 2H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 6.65 (d, 2H), 4.45 (d, 2H), 3.5 (t, 4H), 2.95 (s, 6H). MS(EI) for $\text{C}_{30}\text{H}_{31}\text{N}_7\text{O}_3$: 538 (MH^+).
- [1046] 3-Amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[(3S)-1-(pyridin-2-ylmethyl)pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.93 (s, 1H), 8.65 (d, 1H), 8.56 (s, 1H), 8.48 (m, 1H), 8.32 (d, 1H), 7.9 (d, 1H), 7.72 (m, 2H), 7.6 (t, 1H), 7.46 (d, 1H), 7.38-7.32 (m, 4H), 7.28-7.22 (m, 2H), 4.55 (d, 2H), 4.44 (m, 1H), 3.75 (s, 2H), 2.86-2.64 (m, 2H), 2.64-2.52 (m, 2H), 2.2 (m, 1H). MS (ESI-LCMS) for $\text{C}_{29}\text{H}_{29}\text{N}_7\text{O}_2$: 508 (MH^+).
- [1047] Methyl N-{[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]threoninate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.17 (m, 1H), 9.00 (s, 1H), 8.64 (m, 1H), 8.52 (s, 1H), 8.19 (m, 1H), 7.93 (m, 1H), 7.74-7.62 (m, 3H), 7.37-7.25 (m,

5H), 5.41 (m, 1H), 4.55-4.50 (m, 3H), 4.32-4.29 (m, 1H), 3.70 (m, 3H), 1.16 (m, 2H). MS (EI) for $C_{24}H_{25}N_5O_5$: 464 (MH^+).

[1048] 3-(Ethylamino)-*N*-[(3*S*)-1-ethylpyrrolidin-3-yl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 12.0-10.0 (s, 1H), 9.25-9.15 (tt, 1H), 9.05 (m, 1H), 9.0 (s, 1H), 8.65-8.55 (m, 1H), 8.3 (t, 1H), 7.9 (d, 1H), 7.55 (q, 1H), 7.35-7.2 (m, 5H), 4.75-4.5 (m, 1H), 4.5 (d, 2H), 3.9-3.05 (m, 9 H), 2.3-2.1 (m, 1H), 1.2 (q, 6H). MS (EI) for $C_{27}H_{32}N_6O_2$: 473 (MH^+).

[1049] 3-Amino-*N*-[(3*S*)-1-ethylpyrrolidin-3-yl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.9 (s, 1H), 8.6 (d, 1H), 8.53 (s, 1H), 8.3 (d, 1H), 7.88 (d, 1H), 7.56 (t, 1H), 7.38-7.2 (m, 5H), 4.55 (d, 2H), 4.35 (m, 1H), 2.65 (m, 2H), 2.55-2.35 (m, 4H), 2.15 (m, 1H), 1.8 (m, 1H), 1.0 (t, 3H). MS (EI) for $C_{25}H_{28}N_6O_2$: 445 (MH^+).

[1050] 3-Amino-6-(3-[[[(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl]phenyl]-*N*-[(3*S*)-1-ethylpyrrolidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.95 (m, 2H), 8.6 (d, 1H), 8.55 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 4.4 (m, 1H), 3.0 (m, 1H), 2.85 (m, 1H), 2.7 (m, 1H), 2.6 (m, 1H), 2.5-2.45 (m, 5H), 2.15 (m, 1H), 2.05 (m, 1H), 1.8 (m, 1H), 1.05 (t, 3H). MS (EI) for $C_{27}H_{30}N_6O_2$: 471 (MH^+).

[1051] 6-(3-[[[(1*S*)-2,3-Dihydro-1*H*-inden-1-ylamino]carbonyl]phenyl]-3-(ethylamino)-*N*-[(3*S*)-1-ethylpyrrolidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.0 (s, 2H), 8.75 (t, 1H), 8.7 (d, 1H), 8.6 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 1H), 5.6 (q, 1H), 4.4 (m, 1H), 3.5 (m, 4H), 3.05 (m, 1H), 2.9 (m, 2H), 2.8-2.4 (m, 5H), 2.2-2.0 (m, 2H), 1.8 (m, 1H), 1.2 (t, 3H), 1.05 (t, 3H). MS (EI) for $C_{29}H_{34}N_6O_2$: 499 (MH^+).

[1052] 3-Amino-6-[3-([[(1*S*,2*R*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino]carbonyl]phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.5-9.2 (m, 2H), 9.0 (m, 2H), 8.7 (s, 1H), 8.55 (d, 1H), 8.4 (d, 1H), 7.95 (d, 1H), 7.55 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 4.6 (m, 1H), 4.55 (m, 1H), 3.3-3.05 (m, 5H), 2.95-2.9 (m, 1H), 2.2 5(m, 1H) and 2.1 (m, 1H). MS (EI) for $C_{25}H_{26}N_6O_3$: 459 (MH^+).

- [1053] 1,1-Dimethylethyl (3S)-3-([3-amino-6-(3-([(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.97 (s, 1H), 8.95 (d, 1H), 8.7 (d, 1H), 8.6 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.3-7.1 (m, 4H), 5.6 (t, 1H), 4.45 (t, 1H), 3.6-3.2 (m, 4H), 3.05-2.8 (m, 2H), 2.5 (m, 1H), 2.2-1.9 (m, 3H), 1.4 (s, 9H). MS(EI) for C₃₀H₃₄N₆O₄: 443 (MH⁺).
- [1054] 1,1-Dimethylethyl (3S)-3-([3-amino-6-[3-([(1R)-1-phenylethyl]amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.98 (s, 1H), 8.95 (d, 1H), 8.78 (t, 1H), 8.57 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.42-7.2 (m, 5H), 5.2 (t, 1H), 4.5 (t, 1H), 3.65-3.25 (m, 4H), 2.2-2.0 (m, 2H), 1.5 (d, 3H), 1.4 (s, 9H). MS(EI) for C₂₉H₃₄N₆O₄: 431 (MH⁺).
- [1055] 1,1-Dimethylethyl (3S)-3-([3-amino-6-[3-(3,4-dihydroisoquinolin-2(1*H*)-ylcarbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.9 (t, 1H), 8.75 (t, 1H), 8.3-8.1 (m, 2H), 7.57 (t, 1H), 7.45 (m, 1H), 7.3-7.1 (m, 4H), 4.82 (t, 1H), 4.65-4.4 (m, 2H), 3.6-3.2 (m, 6H), 2.9 (m, 2H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS(EI) for C₃₀H₃₄N₆O₄: 443 (MH⁺).
- [1056] 1,1-Dimethylethyl (3S)-3-([3-amino-6-(3-([(2-phenylethyl]amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.95 (s, 1H), 8.7 (m, 2H), 8.45 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.5 (t, 1H), 3.6-3.45 (m, 4H), 3.4-3.25 (m, 2H), 2.9 (t, 2H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS(EI) for C₂₉H₃₄N₆O₄: 431 (MH⁺).
- [1057] 1,1-Dimethylethyl (3S)-3-([3-amino-6-[3-([(1S)-1-phenylethyl]amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.98 (s, 1H), 8.95 (d, 1H), 8.75 (t, 1H), 8.57 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.45-7.2 (m, 5H), 5.22 (t, 1H), 4.5 (t, 1H), 3.65-3.45 (m, 2H), 3.3 (m, 2H), 2.2-2.0 (m, 2H), 1.55 (s, 3H), 1.4 (s, 9H). MS(EI) for C₂₉H₃₄N₆O₄: 431 (MH⁺).
- [1058] 3-Amino-6-[3-(3,4-dihydroisoquinolin-2(1*H*)-ylcarbonyl)phenyl]-*N*-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.6 (t, 1H), 9.35 (t, 1H), 9.0-8.9 (m, 2H), 8.4 (d, 1H), 8.2 (s, 1H), 7.57 (t, 1H), 7.42 (d, 1H), 7.35-7.0 (m, 4H), 4.8 (s, 1H), 4.6 (m, 2H), 3.9-3.6 (m, 2H), 3.4-3.1 (m, 4H), 2.9 (t, 2H), 2.25 (t, 1H), 2.05 (t, 1H). MS(EI) for C₂₅H₂₆N₆O₂: 443 (MH⁺).

- [1059] 3-Amino-6-(3-([(2-phenylethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.65 (t, 1H), 9.4 (t, 1H), 9.2 (d, 1H), 9.1 (t, 1H), 8.95 (s, 1H), 8.75 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.35-7.2 (m, 5H), 4.7 (t, 1H), 3.6-3.2 (m, 6H), 2.9 (t, 2H), 2.3 (t, 1H), 2.1 (t, 1H). MS(EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431 (MH^+).
- [1060] 3-Amino-6-[3-([(1*R*)-1-phenylethyl]amino)carbonyl]phenyl]-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.75 (t, 1H), 9.55 (t, 1H), 9.3 (m, 2H), 8.99 (s, 1H), 8.8 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.45 (m, 3H), 7.38-7.2 (m, 3H), 5.25 (t, 1H), 4.7 (t, 1H), 3.6-3.2 (m, 4H), 2.3 (t, 1H), 2.15 (t, 1H), 1.6 (s, 3H). MS(EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431 (MH^+).
- [1061] 3-Amino-6-[3-([(1*S*)-1-phenylethyl]amino)carbonyl]phenyl]-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.7 (t, 1H), 9.45 (t, 1H), 9.25 (m, 2H), 9.0 (s, 1H), 8.78 (s, 1H), 8.4 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.5 (m, 2H), 7.35-7.2 (m, 3H), 5.25 (t, 1H), 4.7 (t, 1H), 3.6-3.2 (m, 4H), 2.3 (t, 1H), 2.1 (t, 1H), 1.6 (s, 3H). MS(EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431 (MH^+).
- [1062] 3-(5-Amino-6-cyanopyrazin-2-yl)-*N*-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.99 (s, 1H), 8.96-8.89 (m, 1H), 8.48-8.43 (m, 1H), 8.13-8.07 (m, 1H), 7.98-7.92 (m, 1H), 7.63-7.48 (m, 3H), 7.32-7.16 (m, 4H), 5.66-5.56 (m, 1H), 3.07-2.96 (m, 1H), 2.94-2.80 (m, 1H), 2.54-2.43 (m, 1H), 2.08-1.95 (m, 1H). MS (EI) for $\text{C}_{21}\text{H}_{17}\text{N}_5\text{O}$: 356 (MH^+).
- [1063] 3-Amino-6-[3-([(1*R*,2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl]phenyl]-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.0 (s, 1H), 8.65-8.6 (m, 3H), 8.32 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.3-7.2 (m, 4H), 5.5 (m, 1H), 4.55 (m, 1H), 4.35 (m, 1H), 3.1 (dd, 1H), 3.0-2.85 (m, 4H), 2.75 (m, 3H), 2.0 (m, 1H), and 1.7 (m, 1H). MS (EI) for $\text{C}_{25}\text{H}_{26}\text{N}_6\text{O}_3$: 459 (MH^+).
- [1064] 3-Amino-*N*-[(3*S*)-1-ethylpyrrolidin-3-yl]-6-[3-([(1*S*,2*R*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.0 (s, 1H), 8.7-8.6 (m, 3H), 8.35 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 5.15 (s, 1H), 4.55 (s, 1H), 4.4 (m, 1H), 3.1 (dd, 1H), 2.95 (ss, 1H), 2.75 (m, 1H), 2.6 (s, 1H), 2.5 (m, 4H), 2.2 (m, 1H), 1.9 (m, 1H), 1.05 (t, 3H). MS (EI) for $\text{C}_{27}\text{H}_{30}\text{N}_6\text{O}_3$: 487 (MH^+).

- [1065] 1,1-Dimethylethyl 3-([3-amino-6-(3-([(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)piperidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.95 (m, 2H), 8.55 (m, 2H), 8.3 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 3.9-3.6 (m, 3H), 3.3 (s, 3H), 3.1-2.85 (m, 4H), 2.0 (m, 1H), 1.85 (m, 1H), 1.7 (m, 2H), 1.3 (s, 9H). MS (EI) for C₃₁H₃₆N₆O₄: 457 (MH⁺).
- [1066] 3-Amino-6-(3-([(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)-*N*-piperidin-3-ylpyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.5-9.2 (m, 2H), 9.05 (m, 1H), 8.95 (s, 1H), 8.85 (t, 1H), 8.55 (d, 1H), 8.35 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.25 (m, 4H), 5.6 (m, 1H), 5.0 (s, 1H), 4.3 (s, 1H), 3.3-3.0 (m, 4H), 2.9 (m, 1H), 2.7 (m, 1H), 2.5 (m, 1H), 2.1 (m, 1H), 1.95-1.65 (m, 4H). MS (EI) for C₂₆H₂₈N₆O₂: 457 (MH⁺).
- [1067] 3-Amino-6-(3-([(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)-*N*-(1-ethylpiperidin-3-yl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.95 (m, 2H), 8.55 (m, 2H), 8.25 (d, 1H), 7.92 (d, 1H), 7.6 (t, 1H), 7.25 (m, 4H), 5.6 (q, 1H), 4.0 (m, 1H), 3.05 (m, 1H), 2.9 (m, 1H), 2.7 (m, 1H), 2.35 (m, 2H), 2.2 (s, 2H), 2.05 (m, 1H), 1.7-1.5 (m, 4H), 1.25 (s, 1H), 1.0 (t, 3H). MS (EI) for C₂₈H₃₂N₆O₂: 485 (MH⁺).
- [1068] 3-Amino-*N*-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-([(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.97 (s, 1H), 8.65 (m, 3H), 8.3 (d, 1H), 7.96 (d, 1H), 7.7 (s, 1H), 7.6 (t, 1H), 7.25 (m, 4H), 5.5 (m, 1H), 5.2 (d, 1H), 4.55 (s, 1H), 4.4 (m, 1H), 3.1 (dd, 1H), 2.95 (s, 1H), 2.7 (m, 2H), 2.55-2.4 (m, 4H), 2.15 (m, 1H), 1.85 (m, 1H), 1.05 (t, 3H). MS (EI) for C₂₇H₃₀N₆O₃: 487 (MH⁺).
- [1069] 3-Amino-6-(3-([(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)-*N*-pyrimidin-4-ylpyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 10.9 (s, 1H), 9.2 (s, 2H), 9.04 (d, 1H), 8.9 (d, 1H), 8.65 (s, 1H), 8.4 (d, 2H), 8.05 (d, 1H), 7.7 (t, 1H), 7.3 (m, 4H), 5.7 (m, 1H), 3.15-2.9 (m, 2H), 2.1 (m, 1H). MS (EI) for C₂₅H₂₁N₇O₂: 452 (MH⁺).
- [1070] 3-Amino-6-[3-([(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl]phenyl]-*N*-[2-(1*H*-imidazol-4-yl)ethyl]pyrazine-2-carboxamide: ¹H NMR 7.9 (d, 1H), 7.5 (t, 1H), 7.4 (s, 1H), 7.15 (m, 4H), 5.45 (m, 1H), 4.45 (t, 1H), 3.1 (m, 1H), 2.9 (m, 3H). MS (EI) for C₂₆H₂₅N₇O₃: 484 (MH⁺).

- [1071] 1,1-Dimethylethyl (3S)-3-[[[3-amino-6-(3-[[[(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino)methyl]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.0 (m, 1H), 8.9 (d, 2H), 8.53 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 3.35 (d, 2H), 3.2 (m, 2H), 3.0 (m, 3H), 2.85 (m, 2H), 2.5 (m, 1H), 2.0 (m, 1H), 1.9 (m, 1H), 1.65 (m, 1H), 1.4 (s, 9H). MS (EI) for C₃₁H₃₆N₆O₄: 457 (MH⁺).
- [1072] 3-Amino-6-(3-[[[(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl]phenyl]-*N*-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.4 (t, 2H), 9.22 (t, 1H), 9.05 (d, 1H), 8.95 (s, 1H), 8.6 (s, 1H), 8.5 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.45 (s, 2H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 3.4 (t, 2H), 3.25 (m, 2H), 3.15-2.8 (m, 4H), 2.6 (m, 1H), 2.45 (m, 2H), 2.1 (m, 1H), 2.0 (m, 1H), 1.7 (m, 1H). MS (EI) for C₂₆H₂₈N₆O₂: 457 (MH⁺).
- [1073] 3-Amino-6-{3-[(2,3-dihydro-1*H*-inden-1-ylamino)carbonyl]phenyl}-*N*-piperidin-3-ylpyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.96 (s, 1H), 8.89 (d, *J* = 8.4 Hz, 1H), 8.72 (d, *J* = 8.0 Hz, 1H), 8.54 (s, 1H), 8.35 (d, *J* = 7.2 Hz, 1H), 7.93 (d, *J* = 8.0 Hz, 1H), 7.60 (t, *J* = 9.6 Hz, 1H), 7.30 (m, 1H), 7.25 (m, 1H), 5.61 (q, *J* = 8 Hz, 1H), 4.20 (m, 1H), 3.29 (m, 1H), 3.01 (m, 2H), 2.89 (m, 1H), 2.03 (m, 1H), 1.90 (m, 1H), 1.74 (m, 2H). MS(EI) for C₂₆H₂₈N₆O₂: 457 (MH⁺).
- [1074] 3-Amino-6-(3-[[[(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl]phenyl]-*N*-[(3R)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.96 (s, 1H), 8.89 (d, *J* = 8.2 Hz, 1H), 8.71 (m, 1H), 8.55 (s, 1H), 8.35 (d, *J* = 7.6 Hz, 1H), 7.93 (d, *J* = 8.0 Hz, 1H), 7.60 (t, *J* = 8.0 Hz, 1H), 7.25 (m, 2H), 5.60 (q, *J* = 8.4 Hz, 1H), 4.2 (m, 1H), 3.29 (m, 1H), 3.01 (m, 2H), 2.89 (m, 1H), 2.03 (m, 1H), 1.90 (m, 1H), 1.72 (m, 1H). MS(EI) for C₂₆H₂₈N₆O₂: 457 (MH⁺).
- [1075] 3-Amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino]carbonyl]phenyl]-*N*-piperidin-3-ylpyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.2 (s, 1H), 8.75 (d, 1H), 8.65 (s, 1H), 8.5 (m, 1H), 8.35 (d, 1H), 8.0 (d, 1H), 7.6 (t, 1H), 7.25 (m, 4H), 5.5 (m, 1H), 4.55 (m, 1H), 4.25 (m, 1H), 3.5-2.7 (m, 7H), 1.9 (m, 2H), 1.7 (m, 2H). MS (EI) for C₂₆H₂₈N₆O₃: 473 (MH⁺).
- [1076] 3-Amino-6-(3-[[[(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl]phenyl]-*N*-[2-(1*H*-imidazol-4-yl)ethyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.05 (t, 2H), 8.9 (d, 2H), 8.55 (s, 1H), 8.35 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.5 (s, 1H),

7.3-7.2 (m, 4H) 5.62 (q, 1H), 3.6 (q, 2H), 3.05-2.85 (m, 5H), 2.0 (m, 1H). MS (EI) for $C_{26}H_{25}N_7O_2$: 468 (MH^+).

[1077] 3-Amino-*N*-azepan-4-yl-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.4 (s, 1H), 9.25 (s, 1H), 9.0 (m, 1H), 8.9 (s, 1H), 8.7 (d, 1H), 8.6 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.3-7.1 (m, 4H), 5.6 (q, 1H), 4.1 (s, 1H), 3.25 (m, 1H), 3.1-2.8 (m, 3H), 2.45 (m, 1H), 2.0-1.7 (m, 8H). MS (EI) for $C_{27}H_{30}N_6O_2$: 471 (MH^+).

[1078] 1,1-Dimethylethyl (3*S*)-3-([(3-amino-6-[3-([(1*R*,2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 9.05 (m, 1H), 9.0 (s, 1H), 8.65 (s, 1H), 8.6 (d, 1H), 8.4 (d, 1H), 7.98 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 5.65 (d, 1H), 4.55 (m, 1H), 3.4 (m, 2H), 3.35 (m, 3H), 3.35-3.0 (m, 3H), 2.9 (d, 1H), 1.9 (m, 1H), 1.6 (m, 1H), 1.4 (m, 9H). MS (EI) for $C_{31}H_{36}N_6O_5$: 473 (MH^+).

[1079] 1,1-Dimethylethyl 4-([(3-amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 8.95 (m, 2H), 8.65-8.55 (dd, 2H), 8.35 (d, 1H), 7.95 (d, 1H), 7.7 (s, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 3.95 (m, 1H), 3.6-3.5 (m, 1H), 3.4-3.1 (m, 3H), 3.05 (m, 1H), 2.9 (m, 1H), 2.05-1.4 (m, 7H), 1.4 (s, 9H). MS (EI) for $C_{32}H_{38}N_6O_4$: 471 (MH^+).

[1080] 1,1-Dimethylethyl (3*S*)-3-([(3-amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 8.91 (s, 1H), 8.87 (d, $J = 8.0$ Hz, 1H), 8.51 (m, 1H), 8.27 (d, $J = 6.4$ Hz, 1H), 7.89 (d, $J = 7.2$ Hz, 1H), 7.56 (t, $J = 8.0$ Hz, 1H), 7.27 (m, 1H), 7.21 (m, 1H), 5.60 (q, $J = 8.8$ Hz, 1H), 2.97 (m, 2H), 2.03 (m, 2H), 1.84 (m, 2H), 1.69 (m, 2H), 1.33 (s, 9H). MS (EI) for $C_{31}H_{36}N_6O_4$: 501 ($M-C_4H_9$).

[1081] 1,1-Dimethylethyl (3*S*)-3-([(3-amino-6-[3-([(1*R*,2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino]piperidine-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 8.98 (s, 1H), 8.62 (s, 1H), 8.58 (m, 2H), 8.28 (d, $J = 6.40$ Hz, 1H), 7.93 (d, $J = 8.0$ Hz, 1H), 7.56 (t, 8.0 Hz, 1H), 7.22 (m, 4H), 5.48 (m, 1H), 4.54 (m, 1H), 3.85 (m, 2H), 3.71 (m, 2H), 3.10 (m, 2H), 2.91 (m, 2H), 1.86 (m, 1H), 1.71 (m, 2H), 1.36 (s, 9H). MS (EI) for $C_{31}H_{36}N_6O_5$: 517 ($MH^+ - C_4H_9$).

- [1082] 3-Amino-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino}carbonyl)phenyl]-*N*-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.1 (t, 1H), 8.95 (s, 1H), 8.65 (d, 2H), 8.5 (d, 1H), 7.95 (d, 1H), 7.55 (t, 1H), 7.25-7.15 (m, 4H), 5.5 (m, 1H), 5.0 (s, 1H), 4.55 (m, 1H), 3.3 (m, 2H), 3.15-2.8 (m, 4H), 2.85 (m, 1H), 1.9 (m, 1H), 1.8 (s, 4H), 1.6 (m, 1H). MS (EI) for C₂₆H₂₈N₆O₃: 473 (MH⁺).
- [1083] 1,1-Dimethylethyl 4-[(3-amino-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino}carbonyl)phenyl]pyrazin-2-yl)carbonyl]amino]azepane-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.95 (s, 1H), 8.7-8.65 (m, 3H), 8.45 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 4.55 (m, 1H), 3.95 (m, 1H), 3.6-3.1 (m, 5H), 2.95 (d, 1H), 2.05-1.6 (m, 6H), 1.4 (s, 9H). MS (EI) for C₃₂H₃₈N₆O₅: 487 (MH⁺).
- [1084] 3-Amino-*N*-azepan-4-yl-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.98 (s, 1H), 8.95 (d, 1H), 8.7 (s, 1H), 8.6 (d, 1H), 8.3 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 4.55 (m, 1H), 4.2 (m, 1H), 3.1 (dd, 1H), 3.0-2.7 (m, 4H), 1.95-1.55 (m, 9H). MS (EI) for C₂₇H₃₀N₆O₃: 487 (MH⁺).
- [1085] 3-Amino-*N*-azepan-4-yl-6-[3-[(3*a*R,8*a*S)-8,8*a*-dihydro-3*aH*-indeno[1,2-*d*][1,3]oxazol-2-yl]phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.4-9.25 (dd, 1H), 8.9 (s, 1H), 8.65 (s, 1H), 8.25 (d, 1H), 7.8 (m, 1H), 7.55 (t, 1H), 7.4 (m, 1H), 7.3 (m, 4H), 5.75 (d, 1H), 5.6 (t, 1H), 4.3 (m, 1H), 3.55 (dd, 1H), 3.3 (s, 1H), 3.2-2.9 (m, 4H), 2.7 (m, 1H), 1.9 (m, 2H), 1.65 (m, 3H). MS (EI) for C₂₇H₂₈N₆O₂: 469 (MH⁺).
- [1086] 1,1-Dimethylethyl (3R)-3-[(3-amino-6-(3-({[(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.0 (m, 1H), 8.9 (d, 2H), 8.55 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 3.35 (m, 3H), 3.2 (m, 1H), 3.0 (m, 2H), 2.9 (m, 1H), 2.5 (m, 3H), 2.0 (m, 1H), 1.9 (m, 1H), 1.675 (m, 1H), 1.4 (s, 9H). MS (EI) for C₃₁H₃₆N₆O₄: 557 (MH⁺).
- [1087] 3-Amino-6-(3-({[(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl}phenyl)-*N*-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.35 (s, 2H), 9.15 (t, 1H), 9.0 (d, 1H), 8.85 (s, 1H), 8.6 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.55 (t,

1H), 7.3-7.15 (m, 4H), 5.6 (q, 1H), 4.9 (s, 1H), 3.4 (t, 2H), 3.15-2.85 (m, 4H), 2.65-2.45 (m, 4H), 2.15-1.9 (m, 2H), 1.7 (m, 1H). MS (EI) for C₂₆H₂₈N₆O₂: 457 (MH⁺).

[1088] 1,1-Dimethylethyl (3R)-3-([3-amino-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino}carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]methyl]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, d₆-DMSO): δ 9.0 (m, 1H), 8.95 (s, 1H), 8.65 (s, 1H), 8.57 (d, 1H), 8.4 (d, 1H), 7.95 (d, 1H), 7.55 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 5.1 (d, 1H), 4.55 (m, 1H), 3.35 (m, 5H), 3.25-3.05 (m, 3H), 2.9 (d, 1H), 2.5 (m, 1H), 1.9 (m, 1H), 1.65 (m, 1H), 1.4 (s, 9H). MS (EI) for C₃₁H₃₆N₆O₅: 473 (MH⁺).

[1089] 3-Amino-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino}carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 9.1 (m, 3H), 9.0 (s, 1H), 8.65 (d, 2H), 8.4 (d, 1H), 7.95 (d, 1H), 7.55 (t, 1H), 7.3-7.15 (m, 4H), 5.5 (m, 1H), 5.4 (s, 1H), 4.55 (t, 1H), 3.55 (s, 1H), 3.4 (m, 2H), 3.3-3.05 (m, 4H), 2.95 (d, 2H), 2.6 (m, 1H), 2.0 (m, 1H), 1.7 (m, 1H). MS (EI) for C₂₆H₂₈N₆O₃: 473 (MH⁺).

[1090] 1,1-Dimethylethyl (3R)-3-([3-amino-6-(3-({[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino)piperidine-1-carboxylate: ¹H NMR (400 MHz, d₆-DMSO): δ 8.91 (s, 1H), 8.87 (d, *J* = 8.0 Hz, 1H), 8.51 (m, 1H), 8.27 (d, *J* = 6.4 Hz, 1H), 7.89 (d, *J* = 7.2 Hz, 1H), 7.56 (t, *J* = 8.0 Hz, 1H), 7.27 (m, 1H), 7.21 (m, 1H), 5.60 (q, *J* = 8.8 Hz, 1H), 2.97 (m, 2H), 2.03 (m, 2H), 1.84 (m, 2H), 1.69 (m, 2H), 1.33 (s, 9H). MS (EI) for C₃₁H₃₆N₆O₄: 501 (M-C₄H₉).

[1091] 1,1-Dimethylethyl (3R)-3-([3-amino-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino}carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]piperidine-1-carboxylate: ¹H NMR (400 MHz, d₆-DMSO): δ 8.91 (s, 2H), 8.72 (d, *J* = 8.8 Hz, 1H), 8.51 (m, 1H), 8.28 (d, *J* = 7.6 Hz, 1H), 7.90 (d, *J* = 7.6 Hz, 1H), 7.56 (t, *J* = 8.0 Hz, 1H), 7.27 (m, 2H), 7.19 (m, 2H), 5.60 (m, 4H), 3.81 (m, 2H), 3.71 (m, 2H), 3.00 (m, 2H), 2.91 (m, 2H), 2.01 (m, 1H), 1.86 (m, 1H), 1.71 (m, 2H), 1.36 (s, 9H). MS (EI) for C₃₁H₃₆N₆O₅: 473 (MH⁺-C₅H₉O₂).

[1092] 3-Amino-6-(3-({[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl}phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 9.02 (m, 1H), 8.91 (s, 1H), 8.87 (s, 1H), 8.85 (m, 1H), 8.73 (d, *J* = 8.4 Hz, 1H), 8.65 (s, 1H), 8.57 (m, 1H), 8.52 (m, 1H), 8.43 (m, 1H), 8.35 (d, *J* = 8.0 Hz, 1H), 7.95 (m, 1H), 7.91 (m,

- 1H), 7.62 (m, 2H), 7.36 (t, 8.4 Hz, 1H), 7.17 (m, 1H), 4.62 (d, $J = 6.0$ Hz, 1H), 3.07 (m, 2H), 2.38 (s, 3H), 1.92 (m, 2H), 1.75 (m, 2H). MS (EI) $C_{23}H_{26}N_8O_2$: 447 (MH^+).
- [1093] 3-Amino-6-[3-({[(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino}carbonyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.01(s, 1H), 8.76 (d, $J = 8.4$ Hz, 1H), 8.67 (m, 1H), 8.62 (d, $J = 8.8$ Hz, 1H), 7.97 (d, $J = 8.0$ Hz, 1H), 7.60 (t, 7.6 Hz, 2H), 7.28 (m, 2H), 7.22 (m, 2H), 5.51 (m, 1H), 5.18 (m, 1H), 4.56 (m, 1H), 4.23 (m, 1H), 3.15 (m, 2H), 2.96 (m, 2H), 2.72 (m, 1H), 1.88 (m, 2H), 1.71 (m, 2H), 1.23 (s, 1H). MS (EI) for $C_{26}H_{28}N_6O_3$: 473 (MH^+).
- [1094] 3-(5-Amino-6-[(1*E*)-*N*-[4-(1-methylethyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, $CDCl_3$): δ 8.40 (m, 1H), 8.0 (m, 2H), 7.80 (m, 2H), 7.25 (m, 1H), 7.2 (m, 4H), 6.9 (m, 4H), 4.8 (s, 2H), 2.6 (s, 3H), 2.0 (s, 1H), 1.2 (m, 6H). MS (EI) for $C_{29}H_{30}N_6O$: 479 (MH^+).
- [1095] 3-[5-Amino-6-[(1*E*)-*N*-1,3-benzothiazol-2-ylethanehydrazonoyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.40 (m, 1H), 8.0 (m, 2H), 7.80 (m, 2H), 7.25 (m, 1H), 7.2 (m, 4H), 6.9 (m, 4H), 4.5 (s, 2H), 2.8 (s, 3H). MS (EI) for $C_{27}H_{23}N_7OS$: 494 (MH^+).
- [1096] 3-[5-Amino-6-[(1*E*)-*N*-(4-methylphenyl)ethanehydrazonoyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.90 (d, 1H), 9.80 (d, 1H), 9.20 (d, 1H), 8.50 (d, 2H), 8.20 (d, 1H), 7.90 (m, 1H), 7.75-7.0 (m, 7H), 4.50 (d, 2H), 2.6 (s, 3H), 1.30 (s, 3H). MS (EI) for $C_{27}H_{26}N_6O$: 451 (MH^+).
- [1097] 3-[5-Amino-6-[(1*E*)-*N*-(4-chlorophenyl)ethanehydrazonoyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.80 (s, 1H), 9.10 (m, 1H), 8.65 (s, 1H), 8.55 (s, 1H), 8.2 (d, 1H), 7.90 (d, 1H), 7.60 (m, 1H), 7.4-7.1 (m, 8H), 4.50 (d, 2H), 3.2 (s, 3H), 2.6 (s, 3H). MS (EI) for $C_{27}H_{23}ClN_6O$: 472 (MH^+).
- [1098] 3-[5-Amino-6-[(1*E*)-*N*-methyl-*N*-phenylethanehydrazonoyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 9.00 (d, 1H), 8.80 (d, 1H), 8.20 (dd, 2H), 7.95 (d, 2H), 7.60 (d, 1H), 7.45-7.0 (m, 6H), 4.50 (d, 2H), 3.2 (s, 3H), 2.6 (s, 3H). MS (EI) for $C_{27}H_{26}N_6O$: 451 (MH^+).
- [1099] 3-[5-Amino-6-[(1*E*)-*N*-(2-hydroxyethyl)ethanehydrazonoyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 8.45 (m,

- 2H), 8.15 (d, 1H), 7.80 (d, 1H), 7.45-7.15 (m, 5H), 4.50 (d, 2H), 3.6 (m, 2H), 3.3 (m, 2H), 2.2 (s, 3H). MS (EI) for $C_{22}H_{24}N_6O_2$: 405 (MH^+).
- [1100] 3-{5-Amino-6-[(1E)-N-morpholin-4-ylethanimidoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 8.8 (s, 1H), 8.65 (m, 1H), 8.2 (m, 1H), 7.8 (m, 1H), 7.4 (m, 1H), 7.2 (m, 4H), 4.8 (s, 2H), 3.8 (m, 4H), 3.2 (m, 2H), 2.8 (m, 2H), 2.6 (s, 3H). MS (EI) for $C_{24}H_{26}N_6O_2$: 431 (MH^+).
- [1101] 4-((2E)-2-{1-[3-Amino-6-(3-[[phenylmethyl]amino]carbonyl)phenyl]pyrazin-2-yl}ethylidene)hydrazino)benzoic acid: 1H NMR (400 MHz, d_6 -DMSO): δ 10.0 (s, 1H), 9.2 (m, 1H), 8.8 (s, 1H), 8.5 (s, 1H), 8.2 (m, 1H), 7.8 (m, 3H), 7.60 (m, 1H), 7.25 (m, 3H), 7.2 (m, 3H), 4.6 (s, 2H), 2.8 (s, 3H). MS (EI) for $C_{27}H_{24}N_6O_3$: 481 (MH^+).
- [1102] Ethyl ((2E)-2-{1-[3-amino-6-(3-[[phenylmethyl]amino]carbonyl)phenyl]pyrazin-2-yl}ethylidene)hydrazino)acetate: 1H NMR (400 MHz, d_4 -MeOH): δ 8.40 (m, 1H), 8.2 (m, 2H), 7.9 (m, 1H), 7.80 (m, 1H), 7.35 (m, 1H), 7.2 (m, 4H), 4.6 (s, 2H), 4.2 (m, 2H), 3.8 (s, 2H), 2.4 (s, 3H), 1.2 (m, 3H). MS (EI) for $C_{24}H_{26}N_6O_3$: 447 (MH^+).
- [1103] 3-{5-Amino-6-[(1E)-N,N-dimethylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, $CDCl_3$): δ 9.2 (s, 1H), 9.0 (s, 1H), 8.40 (m, 1H), 8.2 (m, 1H), 7.80 (m, 2H), 7.25 (m, 1H), 7.2 (m, 3H), 4.8 (s, 2H), 3.3 (s, 6H), 2.6 (s, 3H). MS (EI) for $C_{22}H_{24}N_6O$: 389 (MH^+).
- [1104] 3-(5-Amino-6-{(1E)-N-[hydrazino(imino)methyl]ethanehydrazonoyl}pyrazin-2-yl)-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_4 -MeOH): δ 9.2 (s, 1H), 8.8 (s, 1H), 8.6 (m, 1H), 8.2 (m, 2H), 7.80 (m, 1H), 7.25 (m, 1H), 7.2 (m, 3H), 4.6 (s, 2H), 2.8 (s, 3H). MS (EI) for $C_{21}H_{23}N_9O$: 418 (MH^+).
- [1105] 3-(5-Amino-6-{(1E)-N-[4-(methylsulfonyl)phenyl]ethanehydrazonoyl}pyrazin-2-yl)-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.0 (s, 1H), 9.2 (m, 1H), 8.8 (s, 1H), 8.5 (s, 1H), 8.2 (m, 1H), 7.8 (m, 2H), 7.6 (m, 1H), 7.25 (m, 3H), 7.2 (m, 3H), 4.6 (s, 2H), 3.2 (s, 3H), 2.8 (s, 3H). MS (EI) for $C_{27}H_{26}N_6O_3S$: 515 (MH^+).
- [1106] 3-{5-Amino-6-[(1E)-N-(4-cyanophenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.1 (s, 1H), 9.20 (m, 1H), 8.70 (s, 1H), 8.55 (s, 1H), 8.20 (d, 1H), 7.95 (d, 1H), 7.75 (d, 2H), 7.60 (m, 1H), 7.40-7.20 (m, 8H), 4.50 (d, 2H), 2.6 (s, 3H). MS (EI) for $C_{27}H_{23}N_7O$: 462 (MH^+).

- [1107] 3-{5-Amino-6-[(1E)-*N*-pyridin-2-ylethanehydrazonoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 8.75 (s, 1H), 8.55 (s, 1H), 8.20 (d, 2H), 7.95-7.85 (m, 3H), 7.60 (m, 1H), 7.40-7.20 (m, 5H), 4.50 (d, 2H), 2.6 (s, 3H). MS (EI) for $\text{C}_{25}\text{H}_{23}\text{N}_7\text{O}$: 438 (MH^+).
- [1108] 3-(5-Amino-6-[(1E)-*N*-[amino(imino)methyl]ethanehydrazonoyl]pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.0 (m, 1H), 9.20 (m, 1H), 8.80 (s, 1H), 8.75 (s, 1H), 8.20 (d, 2H), 8.00-7.85 (m, 5H), 7.60 (m, 1H), 7.40-7.20 (m, 4H), 4.50 (d, 2H), 2.6 (s, 3H). MS (EI) for $\text{C}_{21}\text{H}_{22}\text{N}_8\text{O}$: 403 (MH^+).
- [1109] 3-[5-Amino-6-[(1E)-*N*-(4-[(trifluoromethyl)oxy]phenyl)ethanehydrazonoyl]pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.1 (s, 1H), 9.90 (s, 1H), 9.20 (m, 1H), 8.70 (s, 1H), 8.55 (s, 1H), 8.20 (d, 1H), 7.95 (d, 2H), 7.60 (m, 2H), 7.40-7.20 (m, 6H), 4.50 (d, 2H), 2.6 (s, 3H). MS (EI) for $\text{C}_{27}\text{H}_{23}\text{F}_3\text{N}_6\text{O}_2$: 522 (MH^+).
- [1110] 3-{5-Amino-6-[(1E)-*N*-(4-nitrophenyl)ethanehydrazonoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.5 (s, 1H), 9.20 (m, 1H), 8.70 (s, 1H), 8.55 (s, 1H), 8.20 (m, 3H), 7.95 (d, 2H), 7.60 (m, 1H), 7.40-7.20 (m, 7H), 4.50 (d, 2H), 2.6 (s, 3H). MS (EI) for $\text{C}_{26}\text{H}_{25}\text{N}_7\text{O}_3$: 483 (MH^+).
- [1111] 3-(5-Amino-6-[(1E)-*N*-[4-(trifluoromethyl)pyrimidin-2-yl]ethanehydrazonoyl]pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.95 (m, 1H), 8.55 (s, 1H), 8.40 (s, 1H), 8.20 (d, 1H), 7.95 (d, 1H), 7.60 (m, 1H), 7.40-7.20 (m, 5H), 4.50 (s, 2H), 2.6 (s, 3H). MS (EI) for $\text{C}_{25}\text{H}_{21}\text{F}_3\text{N}_8\text{O}$: 507 (MH^+).
- [1112] 3-(5-Amino-6-[(1E)-*N*-[(dimethylamino)carbonothioyl]ethanehydrazonoyl]pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.90 (s, 1H), 9.20 (m, 1H), 8.70 (s, 1H), 8.55 (s, 1H), 8.20 (d, 1H), 7.95 (d, 1H), 7.60 (m, 1H), 7.40-7.20 (m, 6H), 4.50 (d, 2H), 3.4 (s, 6H), 2.6 (s, 3H). MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_7\text{OS}$: 448 (MH^+).
- [1113] 3-{5-Amino-6-[(1E)-*N*-1*H*-1,2,3-benzotriazol-1-ylethanimidoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 9.00 (s, 1H), 8.60 (s, 1H), 8.20 (d, 1H), 8.10 (s, 2H), 7.90 (dd, 2H), 7.70-7.60 (m, 2H), 7.50 (m, 1H), 7.40-7.20 (m, 6H), 4.50 (d, 2H), 2.95 (s, 3H). MS (EI) for $\text{C}_{26}\text{H}_{22}\text{N}_8\text{O}$: 463 (MH^+).

- [1114] 3-{5-Amino-6-[(1E)-*N*-methylethanehydrazonoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.50 (d, 1H), 8.30 (s, 1H), 8.20 (d, 1H), 7.90 (d, 1H), 7.65 (m, 1H), 7.40-7.20 (m, 5H), 4.50 (d, 2H), 2.40 (s, 3H), 1.95 (s, 3H). MS (EI) for $\text{C}_{21}\text{H}_{22}\text{N}_6\text{O}$: 375 (MH^+).
- [1115] 3-(5-Amino-6-{(1E)-*N*-[4-(trifluoromethyl)phenyl]ethanehydrazonoyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.0 (s, 1H), 9.2 (m, 1H), 8.8 (s, 1H), 8.5 (s, 1H), 8.2 (m, 1H), 7.8 (m, 2H), 7.6 (m, 1H), 7.25 (m, 3H), 7.2 (m, 3H), 4.6 (s, 2H), 2.8 (s, 3H). MS (EI) for $\text{C}_{27}\text{H}_{23}\text{N}_6\text{OF}_3$: 505 (MH^+).
- [1116] 3-{5-Amino-6-[(1E)-*N*-phenylethanehydrazonoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.0 (s, 1H), 9.2 (m, 1H), 8.8 (s, 1H), 8.5 (s, 1H), 8.2 (m, 1H), 7.8 (m, 2H), 7.6 (m, 1H), 7.25 (m, 4H), 7.2 (m, 3H), 4.6 (s, 2H), 2.8 (s, 3H). MS (EI) for $\text{C}_{26}\text{H}_{24}\text{N}_6\text{O}$: 437 (MH^+).
- [1117] 3-{5-Amino-6-[(1E)-*N*-(4-methylpiperazin-1-yl)ethanimidoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 8.8 (s, 1H), 8.65 (m, 1H), 8.2 (m, 1H), 7.8 (m, 1H), 7.4 (m, 1H), 7.2 (m, 4H), 4.8 (s, 2H), 3.6 (m, 2H), 3.4 (m, 2H), 3.2 (m, 2H), 3.0 (m, 2H), 2.8 (s, 3H), 2.6 (s, 3H). MS (EI) for $\text{C}_{25}\text{H}_{29}\text{N}_7\text{O}$: 444 (MH^+).
- [1118] 3-[5-Amino-6-((1E)-*N*-{(2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl}ethanimidoyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.65 (m, 1H), 8.3 (m, 1H), 8.2 (m, 1H), 7.8 (m, 1H), 7.6 (m, 1H), 7.4 (m, 4H), 7.2 (m, 1H), 4.8 (s, 2H), 3.6 (m, 2H), 3.4 (s, 3H), 3.2 (m, 2H), 3.1 (m, 1H), 2.6 (s, 3H), 2.0 (m, 4H). MS (EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}_2$: 459 (MH^+).
- [1119] 3-{5-Amino-6-[(1E)-*N*-azepan-1-ylethanimidoyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.65 (m, 1H), 8.3 (m, 1H), 8.2 (m, 1H), 7.8 (m, 1H), 7.6 (m, 1H), 7.4 (m, 4H), 7.2 (m, 1H), 4.8 (s, 2H), 3.8 (m, 4H), 2.6 (s, 3H), 2.2 (m, 4H), 1.8 (m, 4H). MS (EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}$: 443 (MH^+).
- [1120] 3-(5-Amino-6-{(E)-[(phenylmethyl)hydrazono]methyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 8.55 (s, 1H), 8.40 (d, 1H), 8.30 (m, 1H), 8.10 (d, 1H), 7.80 (m, 2H), 7.60-7.20 (m, 13H), 4.50-4.40 (dd, 4H). MS (EI) for $\text{C}_{26}\text{H}_{24}\text{N}_6\text{O}$: 437 (MH^+).

- [1121] 3-[5-Amino-6-((E)-{[amino(imino)methyl]hydrazono}methyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.7 (s, 1H), 9.20 (m, 1H), 8.80 (s, 1H), 8.50 (m, 1H), 8.40 (m, 2H), 8.15 (d, 1H), 7.95 (m, 3H), 7.60 (m, 1H), 7.40-7.20 (m, 6H), 4.50 (d, 2H). MS (EI) for $\text{C}_{20}\text{H}_{20}\text{N}_8\text{O}$: 389 (MH^+).
- [1122] 3-(5-Amino-6-{(E)-[(2-hydroxyethyl)hydrazono]methyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (m, 1H), 8.50 (m, 2H), 8.20 (d, 1H), 7.90 (m, 2H), 7.40-7.20 (m, 5H), 4.50 (d, 2H), 3.60 (m, 2H), 3.30 (m, 2H). MS (EI) for $\text{C}_{21}\text{H}_{22}\text{N}_6\text{O}_2$: 391 (MH^+).
- [1123] 3-{5-Amino-6-[(E)-(pyridin-2-ylhydrazono)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.3 (s, 1H), 9.20 (m, 1H), 8.85 (s, 1H), 8.50 (s, 1H), 8.35 (s, 1H), 8.20 (dd, 2H), 7.75-7.50 (m, 3H), 7.40-7.20 (m, 5H), 7.10 (d, 1H), 6.95 (m, 1H), 4.50 (d, 2H). MS (EI) for $\text{C}_{24}\text{H}_{21}\text{N}_7\text{O}$: 424 (MH^+).
- [1124] 3-(5-Amino-6-{(E)-[(4-cyanophenyl)hydrazono]methyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.5 (s, 1H), 9.20 (m, 1H), 8.70 (s, 1H), 8.50 (s, 1H), 8.35 (s, 1H), 8.20 (d, 1H), 7.95 (d, 1H), 7.80-7.55 (m, 5H), 7.40-7.00 (m, 7H), 4.50 (d, 2H). MS (EI) for $\text{C}_{26}\text{H}_{21}\text{N}_7\text{O}$: 448 (MH^+).
- [1125] 3-(5-Amino-6-{(E)-[(4-methylphenyl)hydrazono]methyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.8 (s, 1H), 9.20 (m, 1H), 8.60 (s, 1H), 8.50 (s, 1H), 8.20 (m, 2H), 7.95 (d, 1H), 7.50 (m, 2H), 7.40-6.90 (m, 6H), 4.50 (d, 2H), 2.10 (s, 3H). MS (EI) for $\text{C}_{26}\text{H}_{42}\text{N}_6\text{O}$: 437 (MH^+).
- [1126] 3-{5-Amino-6-[(E)-(hydroxyimino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.9 (s, 1H), 9.20 (s, 1H), 8.75 (s, 1H), 8.55 (s, 1H), 8.30 (s, 1H), 8.10 (d, 1H), 7.90 (s, 1H), 7.65-7.15 (m, 7H), 4.50 (d, 2H). MS (EI) for $\text{C}_{19}\text{H}_{17}\text{N}_5\text{O}_2$: 348 (MH^+).
- [1127] Ethyl 1-{[3-amino-6-(3-{[(phenylmethyl)amino]carbonyl}phenyl)pyrazin-2-yl]methyl}piperidine-4-carboxylate: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.63 (s, 1H), 8.48 (s, 1H), 8.14 (m, 1H), 7.84 (m, 1H), 7.57 (t, 1H), 7.31-7.38 (m, 4H), 7.23-7.27 (m, 1H), 4.61 (s, 2H), 4.43 (s, 2H), 4.16-4.23 (m, 2H), 3.88-3.91 (m, 1H), 3.41-3.68 (m, 1H), 3.29-3.31 (m, 5H), 2.65-2.83 (m, 1H), 2.01-2.32 (m, 4H). MS (EI) for $\text{C}_{27}\text{H}_{31}\text{N}_5\text{O}_3$: 474 (MH^+).

- [1128] 3-[5-Amino-6-(hydroxymethyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.40-8.47 (m, 1H), 8.34 (s, 1H), 8.11-8.14 (m, 1H), 7.86-7.88 (m, 1H), 7.56 (m, 1H), 7.31-7.38 (m, 4H) 7.25-7.26 (m, 1H), 4.82 (s, 2H), 4.61 (s, 2H). MS (EI) for $\text{C}_{19}\text{H}_{18}\text{N}_4\text{O}_2$: 432 (MH^+).
- [1129] 3-{5-Amino-6-[(4-pyridin-2-ylpiperazin-1-yl)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.62 (s, 1H), 8.56 (s, 1H), 8.07-8.16 (m, 3H), 7.58-7.87 (m, 1H), 7.56 (t, 1H), 7.35-7.41 (m, 3H), 7.26-7.30 (m, 2H), 7.16-7.19 (m, 2H), 4.61 (s, 2H), 4.53 (s, 2H), 4.05 (s, 4H), 3.71-3.73 (m, 4H). MS (EI) for $\text{C}_{28}\text{H}_{29}\text{N}_7\text{O}$: 480 (MH^+).
- [1130] 3-(5-Amino-6-{[3-(hydroxymethyl)piperidin-1-yl]methyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.63 (s, 1H), 8.51 (t, 1H), 8.13-8.15 (m, 1H), 7.85-7.87 (m, 1H), 7.55-7.59 (m, 1H), 7.32-7.39 (m, 4H), 7.20-7.28 (m, 1H), 4.59-4.70 (m, 2H), 4.41 (s, 2H), 4.05-4.08 (m, 1H), 3.75-3.78 (m, 1H), 3.51-3.58 (m, 1H), 3.34-3.40 (m, 1H), 3.12-3.17 (m, 1H), 2.89 (t, 1H), 1.77-2.16 (m, 4H), 1.30-1.36 (m, 1H). MS (ESI-LCMS) for $\text{C}_{25}\text{H}_{29}\text{N}_5\text{O}_2$: 432 (MH^+).
- [1131] 1,1-Dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}pyrazin-2-yl)carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.06 (t, 1H), 8.95 (s, 1H), 8.7 (d, 1H), 8.5 (s, 1H), 8.3 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.25 (d, 2H), 6.85 (d, 2H), 4.5 (d, 2H), 3.7 (s, 3H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{29}\text{H}_{34}\text{N}_6\text{O}_5$: 447 (MH^+).
- [1132] 1,1-Dimethylethyl (3S)-3-[[[3-amino-6-(3-[[[4-(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.23 (t, 1H), 8.95 (s, 1H), 8.75 (d, 1H), 8.55 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.48 (d, 2H), 7.37 (d, 2H), 4.57 (d, 2H), 4.5 (m, 1H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{29}\text{H}_{31}\text{N}_6\text{O}_5\text{F}_3$: 501 (MH^+).
- [1133] 1,1-Dimethylethyl (3S)-3-[[[3-amino-6-[3-[[[2-chloro-6-fluorophenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.9 (s, 1H), 8.8 (d, 1H), 8.7 (t, 1H), 8.45 (s, 1H), 8.25 (d, 1H), 7.8 (d, 1H), 7.5 (t, 1H), 7.35-7.15 (m, 3H), 4.6 (d, 2H), 4.4 (m, 1H), 3.6-3.2 (m, 4H), -2.1-1.9 (m, 2H), 1.35 (s, 9H). MS (EI) for $\text{C}_{28}\text{H}_{30}\text{N}_6\text{O}_4\text{Cl}$: 469 (MH^+).

- [1134] 1,1-Dimethylethyl (3S)-3-[[[(3-amino-6-{3-[[[4-(methylthio)phenyl]methyl]amino)carbonyl]phenyl}pyrazin-2-yl)carbonyl]amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.1 (t, 1H), 8.9 (s, 1H), 8.7 (d, 1H), 8.5 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.3 (d, 2H), 7.2 (d, 2H), 4.6-4.4 (m, 3H), 3.6-3.2 (m, 4H), 2.4 (s, 3H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for C₂₉H₃₄N₆O₄S: 463 (MH⁺).
- [1135] 1,1-Dimethylethyl (3S)-3-[[[(3-amino-6-{3-[(2,3-dihydro-1*H*-inden-2-ylamino)carbonyl]phenyl}pyrazin-2-yl)carbonyl]amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.95 (s, 1H), 8.8-8.7 (m, 2H), 8.5 (s, 1H), 8.3 (d, 1H), 7.83 (d, 1H), 7.57 (t, 1H), 7.2 (m, 2H), 7.15 (m, 2H), 4.7 (m, 2H), 3.6-3.4 (m, 2H), 3.25 (m, 4H), 2.98 (m, 2H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for C₃₀H₃₄N₆O₄: 443 (MH⁺).
- [1136] 3-Amino-6-{3-[[[4-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-*N*-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.6-9.4 (m, 3H), 9.2 (d, 1H), 8.95 (s, 1H), 8.7 (s, 1H), 8.5 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.3 (d, 2H), 6.85 (d, 2H), 4.7 (m, 1H), 4.4 (d, 2H), 3.7 (s, 3H), 3.4-3.1 (m, 4H), 2.25 (t, 1H), 2.05 (t, 1H). MS (EI) for C₂₄H₂₆N₆O₃: 447 (MH⁺).
- [1137] 3-Amino-*N*-[(3S)-pyrrolidin-3-yl]-6-(3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.65 (t, 1H), 9.5 (m, 1H), 9.25 (m, 1H), 9.2 (d, 1H), 8.95 (s, 1H), 8.75 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.5 (d, 2H), 7.3 (d, 2H), 4.7 (t, 1H), 4.5 (d, 2H), 3.4-3.1 (m, 4H), 2.25 (t, 1H), 2.05 (t, 1H). MS (EI) for C₂₄H₂₃N₆O₃F₃: 501 (MH⁺).
- [1138] 3-Amino-6-{3-[(2,3-dihydro-1*H*-inden-2-ylamino)carbonyl]phenyl}-*N*-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.65 (m, 1H), 9.45 (m, 1H), 9.2 (m, 2H), 8.96 (s, 1H), 8.72 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.25 (t, 2H), 7.18 (d, 2H), 4.8-4.6 (m, 2H), 3.45-3.1 (m, 8H), 2.3 (t, 1H), 2.1 (t, 1H). MS (EI) for C₂₅H₂₆N₆O₂: 443 (MH⁺).
- [1139] 3-Amino-6-[3-([[(2-chloro-6-fluorophenyl)methyl]amino]carbonyl]phenyl]-*N*-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.5 (m, 1H), 9.35 (m, 1H), 9.1 (m, 2H), 8.95 (s, 1H), 8.6 (s, 1H), 8.4 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.4-7.2 (m, 3H), 4.65 (s, 3H), 3.4-3.2 (m, 4H), 2.3 (m, 1H), 2.05 (m, 1H). MS (EI) for C₂₃H₂₂N₆O₂FCI: 469 (MH⁺).

- [1140] 3-Amino-6-[3-[[[4-(methylthio)phenyl]methyl]amino]carbonyl]phenyl]-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.25 (t, 1H), 8.9 (s, 1H), 8.86 (d, 1H), 8.6 (s, 1H), 8.3 (d, 1H), 7.85 (d, 1H), 7.62 (d, 1H), 7.57 (t, 1H), 7.3 (d, 2H), 7.2 (d, 2H), 4.6 (t, 1H), 4.45 (d, 2H), 3.4-3.1 (m, 4H), 2.4 (s, 3H), 2.2 (m, 1H), 2.0 (m, 1H). MS (EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2\text{S}$: 463 (MH^+).
- [1141] 1,1-Dimethylethyl (3*S*)-3-[[[3-amino-6-[3-[[[4-chlorophenyl]methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.95 (s, 1H), 8.75 (d, 1H), 8.55 (s, 1H), 8.4 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.4 (s, 4H), 4.5 (d, 2H), 4.47 (m, 1H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{28}\text{H}_{31}\text{N}_6\text{O}_4\text{Cl}$: 451 (MH^+).
- [1142] 1,1-Dimethylethyl (3*S*)-3-[[[3-amino-6-(3-[[[pyridin-2-ylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.4 (t, 1H), 8.95 (s, 1H), 8.7 (d, 2H), 8.6 (s, 1H), 8.4 (d, 1H), 8.2 (m, 1H), 7.9 (d, 1H), 7.75 (d, 1H), 7.65 (t, 1H), 7.6 (m, 2H), 4.75 (d, 2H), 4.5 (m, 1H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{27}\text{H}_{31}\text{N}_7\text{O}_4$: 462 (MH^+).
- [1143] 1,1-Dimethylethyl (3*S*)-3-[[[3-amino-6-(3-[[[pyridin-3-ylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (t, 1H), 8.9 (s, 1H), 8.85 (m, 1H), 8.78 (d, 1H), 8.7 (d, 1H), 8.5 (s, 1H), 8.43 (d, 1H), 8.38 (d, 1H), 7.9 (m, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 4.7 (d, 2H), 4.5 (m, 1H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{27}\text{H}_{31}\text{N}_7\text{O}_4$: 518 (MH^+).
- [1144] 1,1-Dimethylethyl (3*S*)-3-[[[3-amino-6-[3-[(3,4-dihydro-2*H*-chromen-4-ylamino)carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.0 (m, 1H), 8.9 (s, 1H), 8.55 (s, 1H), 8.38 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.22 (d, 1H), 7.17 (t, 1H), 6.85 (t, 1H), 6.78 (d, 1H), 5.3 (m, 1H), 4.6-4.4 (m, 1H), 4.4-4.2 (m, 2H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 4H), 1.4 (s, 9H). MS (EI) for $\text{C}_{30}\text{H}_{34}\text{N}_6\text{O}_5$: 459 (MH^+).
- [1145] 3-Amino-6-[3-[[[4-chlorophenyl]methyl]amino]carbonyl]phenyl]-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.55 (t, 1H), 9.37 (m, 1H), 9.1 (m, 2H), 8.95 (s, 1H), 8.7 (s, 1H), 8.5 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.4 (s, 4H), 4.7 (m, 1H), 4.5 (d, 2H), 3.5-3.3 (m, 3H), 3.3-3.1 (m, 1H), 2.3 (m, 1H), 2.1 (m, 1H). MS (EI) for $\text{C}_{23}\text{H}_{23}\text{N}_6\text{O}_2\text{Cl}$: 451 (MH^+).

- [1146] 3-Amino-6-(3-([(pyridin-2-ylmethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.15 (t, 1H), 9.55-9.4 (m, 2H), 9.26 (d, 1H), 8.95 (m, 2H), 8.82 (d, 1H), 8.5 (m, 1H), 8.35 (d, 1H), 8.0 (d, 1H), 7.9 (m, 2H), 7.57 (t, 1H), 4.9 (d, 2H), 4.75 (m, 1H), 3.4 (m, 3H), 3.2 (m, 1H), 2.3 (m, 1H), 2.1 (m, 1H). MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_7\text{O}_2$: 418 (MH^+).
- [1147] 3-Amino-6-(3-([(pyridin-3-ylmethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.15 (t, 1H), 9.6 (m, 2H), 9.3 (d, 1H), 9.03-8.83 (m, 4H), 8.65 (d, 1H), 8.3 (d, 1H), 8.07 (t, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 4.78 (m, 3H), 3.42 (m, 3H), 3.2 (m, 1H), 2.3 (m, 1H), 2.1 (m, 1H). MS (EI) for $\text{C}_{22}\text{H}_{23}\text{N}_7\text{O}_2$: 418 (MH^+).
- [1148] 3-Amino-6-{3-[(3,4-dihydro-2H-chromen-4-ylamino)carbonyl]phenyl}-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.0 (m, 2H), 8.95-8.75 (m, 3H), 8.57 (s, 1H), 8.37 (d, 1H), 7.95 (d, 1H), 7.6 (t, 1H), 7.2 (m, 2H), 6.9 (t, 1H), 6.82 (d, 1H), 5.35 (m, 1H), 4.6 (m, 1H), 4.3 (m, 2H), 3.4 (m, 2H), 3.25 (m, 2H), 2.3-2.0 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{26}\text{N}_6\text{O}_3$: 459 (MH^+).
- [1149] 3-Amino-6-{3-[(2,3-dihydro-1-benzofuran-3-ylamino)carbonyl]phenyl}-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (d, 1H), 8.99 (m, 2H), 8.88 (d, 1H), 8.6 (s, 1H), 8.38 (d, 1H), 7.95 (d, 1H), 7.57 (t, 1H), 7.4 (d, 1H), 7.26 (t, 1H), 6.95 (m, 2H), 5.85 (m, 1H), 4.8-4.5 (m, 3H), 4.4-4.0 (m, 2H), 3.5-3.2 (m, 2H), 2.4-2.0 (m, 2H). MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_3$: 445 (MH^+).
- [1150] 1,1-Dimethylethyl (3*S*)-3-([(3-amino-6-(3-([(1-methyl-1-phenylethyl)amino]carbonyl)phenyl)pyrazin-2-yl]carbonyl)amino)pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.95 (s, 1H), 8.75 (m, 1H), 8.75 (s, 1H), 8.3 (d, 1H), 7.75 (d, 1H), 7.55 (t, 1H), 7.4 (d, 2H), 7.25 (t, 2H), 7.15 (m, 1H), 4.5 (m, 1H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.7 (s, 6H), 1.4 (s, 9H). MS (EI) for $\text{C}_{30}\text{H}_{36}\text{N}_6\text{O}_4$: 445 (MH^+).
- [1151] 3-Amino-6-(3-([(1-methyl-1-phenylethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.1-8.9 (m, 2H), 9.0 (s, 1H), 8.95 (d, 1H), 8.6 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 7.4 (d, 2H), 7.3 (t, 2H), 7.2 (m, 1H), 4.6 (m, 1H), 3.5-3.2 (m, 4H), 2.3-2.1 (m, 2H), 1.7 (s, 6H). MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2$: 445 (MH^+).

- [1152] 3-Amino-6-(3-([(1S)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)-*N*-[(3S)-1-methylpyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.81-8.75 (m, 1H), 8.59-8.54 (m, 1H), 8.26-8.18 (m, 1H), 7.90-7.85 (m, 1H), 7.61-7.54 (m, 1H), 7.34-7.16 (m, 4H), 5.73-5.66 (m, 1H), 4.71-4.61 (m, 1H), 4.05-3.85 (m, 2H), 3.50-3.41 (m, 1H), 3.29-2.87 (m, 6H), 2.76-2.56 (m, 2H), 2.44-2.23 (m, 1H), 2.09-2.01 (m, 1H). MS (EI) for C₂₆H₂₈N₆O₂: 457 (MH⁺).
- [1153] 3-Amino-6-[3-([(1R,2S)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)carbonyl]phenyl)-*N*-[(3S)-1-methylpyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.77-8.71 (m, 1H), 8.64-8.59 (m, 1H), 8.25-8.19 (m, 1H), 8.00-7.93 (m, 1H), 7.64-7.57 (m, 1H), 7.35-7.18 (m, 4H), 5.64-5.59 (m, 1H), 4.79-4.60 (m, 2H), 3.96-3.84 (m, 1H), 3.49-3.40 (m, 1H), 3.28-3.13 (m, 2H), 3.05-2.97 (m, 5H), 2.73-2.61 (m, 1H), 2.41-2.23 (m, 1H). MS (EI) for C₂₆H₂₈N₆O₃: 473 (MH⁺).
- [1154] 3-Amino-*N*-[(3S)-pyrrolidin-3-yl]-6-[3-([(3,4,5-trifluorophenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.8 (t, 1H), 9.5 (s, 1H), 9.3 (s, 1H), 9.2 (d, 1H), 8.95 (s, 1H), 8.8 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.35 (m, 2H), 4.75 (m, 1H), 4.5 (d, 2H), 3.4 (m, 3H), 3.2 (m, 1H), 2.3 (m, 1H), 2.15 (m, 1H). MS (EI) for C₂₃H₂₁N₆O₂F₃: 471 (MH⁺).
- [1155] 3-Amino-*N*-[(3S)-pyrrolidin-3-yl]-6-[3-([(2,3,6-trifluorophenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.55 (s, 1H), 9.45 (s, 1H), 9.35 (t, 1H), 9.1 (d, 1H), 8.9 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.55 (t, 1H), 7.45 (m, 1H), 7.12 (m, 1H), 5.0 (s, 1H), 4.65 (m, 1H), 4.55 (d, 2H), 3.45-3.15 (m, 4H), 2.25 (m, 1H), 2.1 (m, 1H). MS (EI) for C₂₃H₂₁N₆O₂F₃: 471 (MH⁺).
- [1156] 3-Amino-*N*-[(3S)-pyrrolidin-3-yl]-6-[3-([(2,3,5-trifluorophenyl)methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.8 (t, 1H), 9.55 (s, 1H), 9.4 (s, 1H), 9.25 (d, 1H), 8.95 (s, 1H), 8.85 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.55 (t, 1H), 7.45 (m, 1H), 7.15 (m, 1H), 4.75 (m, 1H), 4.55 (d, 2H), 3.4 (m, 3H), 3.2 (m, 1H), 2.3 (m, 1H), 2.15 (m, 1H). MS (EI) for C₂₃H₂₁N₆O₂F₃: 471 (MH⁺).
- [1157] 3-Amino-6-[3-([(3-bromo-4-fluorophenyl)methyl]amino)carbonyl]phenyl]-*N*-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.7 (t, 1H), 9.5 (s, 1H), 9.3 (s, 1H), 9.2 (d, 1H), 8.9 (s, 1H), 8.75 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.7 (d, 1H), 7.55 (t, 1H), 7.4 (m, 1H), 7.3 (t, 1H), 4.7 (m, 1H), 4.5 (d, 2H), 3.4 (m, 3H), 3.2 (m, 1H), 2.3 (m, 1H), 2.15 (m, 1H). MS (EI) for C₂₃H₂₂N₆O₂FB: 515 (MH⁺).

- [1158] 1,1-Dimethylethyl (3S)-3-[(3-amino-6-[3-((2,4-dichlorophenyl)methyl)amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.16 (t, 1H), 8.9 (s, 1H), 8.7 (d, 1H), 8.55 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.6 (s, 1H), 7.57 (t, 1H), 7.4 (m, 2H), 4.58 (d, 2H), 4.5 (m, 1H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{28}\text{H}_{30}\text{N}_6\text{O}_4\text{Cl}_2$: 485 (MH^+).
- [1159] 1,1-Dimethylethyl (3S)-3-[(3-amino-6-[3-((3,4-dichlorophenyl)methyl)amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.9 (s, 1H), 8.7 (d, 1H), 8.5 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.6 (m, 3H), 7.36 (d, 1H), 4.5 (m, 3H), 3.6-3.2 (m, 4H), 2.2-2.0 (m, 2H), 1.4 (s, 9H). MS (EI) for $\text{C}_{28}\text{H}_{30}\text{N}_6\text{O}_4\text{Cl}_2$: 485 (MH^+).
- [1160] 3-Amino-6-[3-((2,4-dichlorophenyl)methyl)amino]carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.7 (t, 1H), 9.5 (m, 1H), 9.3 (m, 1H), 9.2 (d, 1H), 8.95 (s, 1H), 8.8 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (s, 1H), 7.57 (t, 1H), 7.4 (d, 2H), 4.7 (m, 1H), 4.55 (d, 2H), 3.5-3.1 (m, 4H), 2.4-2.0 (m, 2H). MS (EI) for $\text{C}_{23}\text{H}_{22}\text{N}_6\text{O}_2\text{Cl}_2$: 485 (MH^+).
- [1161] 3-Amino-6-[3-((3,4-dichlorophenyl)methyl)amino]carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.78 (t, 1H), 9.5 (m, 1H), 9.35 (m, 1H), 9.23 (d, 1H), 8.95 (s, 1H), 8.8 (s, 1H), 8.53 (d, 1H), 7.9 (d, 1H), 7.6 (s, 1H), 7.6-7.5 (m, 2H), 7.4 (d, 1H), 4.7 (m, 1H), 4.5 (d, 2H), 3.5-3.1 (m, 4H), 2.4-2.1 (m, 2H). MS (EI) for $\text{C}_{23}\text{H}_{22}\text{N}_6\text{O}_2\text{Cl}_2$: 485 (MH^+).
- [1162] 3-Amino-6-[3-((4-(methyloxy)phenyl)methyl)amino]carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.43 (t, 1H), 9.4 (m, 1H), 9.15 (m, 1H), 8.9 (s, 1H), 8.65 (s, 1H), 8.3 (d, 1H), 7.87 (d, 1H), 7.57 (t, 1H), 7.3 (d, 2H), 6.85 (d, 2H), 4.45 (d, 2H), 3.7 (s, 3H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.9-1.7 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_3$: 461 (MH^+).
- [1163] 3-Amino-6-[3-((2,4-dichlorophenyl)methyl)amino]carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.65 (t, 1H), 9.3 (m, 1H), 9.05 (m, 1H), 9.0-8.9 (m, 2H), 8.75 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.6-7.5 (m, 2H), 7.4 (d, 2H), 4.6 (d, 2H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.9-1.7 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2\text{Cl}_2$: 499 (MH^+).
- [1164] 3-Amino-6-[3-((4-fluorophenyl)methyl)amino]carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.4 (t, 1H),

9.05 (d, 1H), 8.95 (s, 1H), 8.85 (d, 2H), 8.65 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.4 (m, 2H), 7.15 (m, 2H), 4.5 (d, 2H), 4.25 (m, 1H), 3.25 (m, 2H), 3.1 (m, 1H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_2F$: 449 (MH^+).

[1165] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.55 (t, 1H), 9.15 (d, 1H), 8.95 (s, 1H), 8.0 (d, 2H), 8.7 (s, 1H), 8.35 (s, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.5 (d, 2H), 7.35 (d, 2H), 4.55 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{25}H_{25}N_6O_3F_3$: 515 (MH^+).

[1166] 3-Amino-6-(3-[[[biphenyl-4-ylmethyl]amino]carbonyl}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.4 (t, 1H), 9.05 (d, 1H), 8.95 (s, 1H), 9.35 (d, 2H), 8.65 (s, 1H), 8.35 (d, 1H), 7.95 (d, 1H), 7.7-7.6 (m, 5H), 7.45 (m, 4H), 7.35 (m, 1H), 4.6 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{30}H_{30}N_6O_2$: 507 (MH^+).

[1167] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-{3-[[[4-(1,2,3-thiadiazol-5-yl)phenyl]methyl]amino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.6 (s, 1H), 9.25 (t, 1H), 8.95 (s, 1H), 8.75 (d, 2H), 8.65 (m, 1H), 8.55 (s, 1H), 8.35 (d, 1H), 8.15 (d, 1H), 7.95 (d, 1H), 7.7 (s, 1H), 7.6 (t, 1H), 7.55 (d, 2H), 4.6 (d, 2H), 4.25 (m, 1H), 3.3 (m, 2H), 3.05 (m, 1H), 2.75 (m, 1H), 1.95-1.65 (m, 4H). MS (EI) for $C_{26}H_{26}N_8O_2S$: 515 (MH^+).

[1168] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-[[[2-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.5 (t, 1H), 9.2 (s, 1H), 8.95 (s, 2H), 8.9 (d, 1H), 8.7 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.5 (d, 1H), 7.4 (m, 3H), 4.6 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{25}H_{25}N_6O_2F_3$: 515 (MH^+).

[1169] 3-Amino-6-[3-[[[2,4-difluorophenyl]methyl]amino]carbonyl}phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.5 (t, 1H), 9.25 (m, 1H), 9.05 (m, 1H), 8.95 (s, 1H), 8.93 (d, 1H), 8.7 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t, 1H), 7.5 (m, 1H), 7.25 (m, 1H), 7.1 (m, 1H), 4.55 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{24}H_{24}N_6O_2F_2$: 467 (MH^+).

[1170] 3-Amino-6-[3-[[[4-chloro-2-fluorophenyl]methyl]amino]carbonyl}phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.5 (t, 1H), 9.15 (d, 1H), 8.95 (s, 1H), 8.9 (d, 2H), 8.7 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.6 (t,

1H), 7.45 (m, 2H), 7.3 (m, 1H), 4.55 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{24}H_{24}N_6O_2FCl$: 483 (MH^+).

[1171] 3-Amino-6-{3-[[[4-fluoro-3-(trifluoromethyl)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.55 (t, 1H), 9.05 (d, 1H), 8.95 (s, 1H), 8.85 (m, 2H), 8.65 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.8 (m, 2H), 7.6 (t, 1H), 7.5 (t, 1H), 4.5 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{25}H_{24}N_6O_2F_4$: 517 (MH^+).

[1172] 3-Amino-6-(3-[[[naphthalen-2-ylmethyl]amino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.55 (t, 1H), 9.3 (m, 1H), 9.1 (m, 1H), 9.0-8.9 (m, 2H), 8.75 (s, 1H), 8.4 (d, 1H), 8.3 (d, 1H), 8.0 (m, 2H), 7.9 (d, 1H), 7.7-7.5 (m, 5H), 5.05 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{28}H_{28}N_6O_2$: 481 (MH^+).

[1173] 3-Amino-6-{3-[[[4-(dimethylamino)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.05 (t, 1H), 8.9 (s, 1H), 8.8 (m, 1H), 8.7 (d, 1H), 8.65 (m, 1H), 8.5 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.25 (d, 2H), 6.85 (d, 2H), 4.4 (d, 2H), 4.25 (m, 1H), 3.4-3.0 (m, 3H), 2.9 (s, 6H), 2.9-2.7 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{26}H_{31}N_7O_2$: 474 (MH^+).

[1174] 3-Amino-6-(3-[[[1,3-benzodioxol-5-ylmethyl]amino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.1 (t, 1H), 8.95 (s, 1H), 8.9 (m, 1H), 8.7 (d, 1H), 8.5 (s, 1H), 8.35 (d, 1H), 7.85 (d, 1H), 7.57 (t, 1H), 6.9 (s, 1H), 6.85 (m, 2H), 5.98 (s, 2H), 4.43 (d, 1H), 4.2 (m, 1H), 3.4-3.2 (m, 2H), 3.05 (q, 1H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{25}H_{26}N_6O_4$: 475 (MH^+).

[1175] 3-Amino-N-[(3S)-piperidin-3-yl]-6-{3-[[[4-(2-thienyl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (t, 1H), 9.0 (m, 1H), 8.95 (s, 1H), 8.8 (m, 1H), 8.78 (d, 1H), 8.6 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.64-7.56 (m, 2H), 7.51 (d, 1H), 7.6 (d, 1H), 7.4 (d, 2H), 7.1 (d, 1H), 4.55 (d, 2H), 4.25 (m, 1H), 3.3-3.0 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{28}H_{28}N_6O_2S$: 513 (MH^+).

[1176] 3-Amino-6-{3-[[[2-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.26 (t, 2H), 9.03 (m, 1H), 8.95 (s, 1H), 8.87 (d, 1H), 8.65 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.57 (t,

- 1H), 7.2 (m, 2H), 7.0 (d, 1H), 6.9 (t, 1H), 4.5 (d, 2H), 4.3 (m, 1H), 3.8 (s, 3H), 3.3-3.0 (m, 3H), 2.74 (m, 1H), 1.9-1.7 (m, 4H). MS (EI) for $C_{25}H_{28}N_6O_3$: 461 (MH^+).
- [1177] 3-Amino-6-[3-([2,4-bis(methyloxy)phenyl]methyl)amino]carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (m, 1H), 9.1 (m, 2H), 8.95 (s, 1H), 8.9 (d, 1H), 8.65 (s, 1H), 8.35 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.1 (d, 1H), 6.6 (s, 1H), 6.5 (d, 1H), 4.5 (d, 2H), 4.3 (m, 1H), 3.8 (s, 6H), 3.3-3.0 (m, 3H), 2.75 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{26}H_{30}N_6O_4$: 491 (MH^+).
- [1178] 3-Amino-6-[3-([(4-chloro-2-methylphenyl)methyl]amino)carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.4 (t, 1H), 9.3 (m, 1H), 9.07 (m, 1H), 8.95 (s, 1H), 8.94 (d, 1H), 8.7 (s, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.3 (d, 1H), 7.25 (s, 1H), 7.2 (d, 1H), 4.45 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 2.4 (s, 3H), 2.0-1.7 (m, 4H). MS (EI) for $C_{25}H_{27}N_6O_2Cl$: 479 (MH^+).
- [1179] 3-Amino-6-[3-([(2,3-difluoro-4-methylphenyl)methyl]amino)carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.5 (t, 1H), 9.3 (m, 1H), 9.05 (m, 1H), 8.95 (s, 1H), 8.94 (d, 1H), 8.7 (s, 1H), 8.34 (d, 1H), 7.9 (d, 1H), 7.57 (t, 1H), 7.17 (m, 1H), 7.04 (m, 1H), 4.55 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.78 (m, 1H), 2.25 (s, 3H), 2.0-1.7 (m, 4H). MS (EI) for $C_{25}H_{26}N_6O_2F_2$: 481 (MH^+).
- [1180] 3-Amino-6-[3-([(4-(methylsulfonyl)phenyl)methyl]amino)carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.79 (s, 1H), 8.55 (m, 1H), 8.23-8.20 (m, 1H), 7.95-7.87 (m, 3H), 7.65-7.57 (m, 3H), 4.72 (s, 2H), 4.31-4.29 (m, 1H), 3.50-3.45 (m, 2H), 3.38-3.31 (m, 2H), 3.13-2.98 (m, 5H), 2.10-1.83 (m, 4H). MS (EI) for $C_{25}H_{28}N_6O_4S$: 510 (MH^+).
- [1181] 3-Amino-6-[3-([(2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino]carbonylphenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.78 (s, 1H), 8.53 (s, 1H), 8.19-8.17 (m, 1H), 7.85-7.83 (m, 1H), 7.58-7.55 (m, 1H), 6.86-6.77 (m, 3H), 4.49 (s, 2H), 4.31-4.28 (m, 1H), 4.20 (s, 4H), 3.51-3.46 (m, 1H), 3.37-3.29 (m, 1H), 3.16-3.10 (m, 1H), 2.99-2.97 (m, 1H), 2.10-1.83 (m, 4H). MS (EI) for $C_{26}H_{28}N_6O_4$: 489 (MH^+).
- [1182] 3-Amino-6-[3-([(4-chlorophenyl)methyl]amino)carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.76 (s, 1H),

8.60 (m, 1H), 8.23-8.21 (m, 1H), 7.92-7.89 (m, 1H), 7.62-7.59 (m, 1H), 7.39-7.32 (m, 4H), 4.60 (s, 2H), 4.31-4.29 (m, 1H), 3.50-3.46 (m, 2H), 3.37-3.30 (m, 2H), 2.10-1.83 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_2Cl$: 465 (MH^+).

[1183] 3-Amino-6-[3-({[(5-chloro-2-fluorophenyl)methyl]amino}carbonyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.79 (s, 1H), 8.61 (m, 1H), 8.23-8.20 (m, 1H), 7.87-7.82 (m, 1H), 7.77-7.03 (m, 5H), 4.82 (s, 2H), 4.39-4.36 (m, 1H), 3.51-3.45 (m, 2H), 3.13-2.98 (m, 2H), 2.10-1.83 (m, 4H). MS (EI) for $C_{24}H_{24}N_6O_2FCl$: 483 (MH^+).

[1184] 3-Amino-6-{3-[{[4-(methylthio)phenyl]methyl]amino}carbonyl]phenyl}-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.26 (m, 1H), 8.95 (s, 1H), 8.79 (d, $J = 8.4$ Hz, 1H), 8.58 (s, 1H), 8.34 (d, $J = 8.0$ Hz, 1H), 7.90 (d, $J = 8.4$ Hz, 1H), 7.94 (t, $J = 8.0$ Hz, 1H), 7.31 (m, 2H), 7.24 (m, 2H), 4.48 (d, $J = 6.0$ Hz, 2H), 4.24 (m, 1H), 3.72 (m, 1H), 3.68 (m, 1H), 3.59 (s, 2H), 3.08 (m, 2H), 2.75 (m, 2H), 2.67 (m, 2H), 1.91 (m, 2H), 1.76 (m, 2H). MS (EI) for $C_{25}H_{28}N_6O_2S$: 477 (MH^+).

[1185] 3-Amino-6-(3-[{(1,2-diphenylethyl)amino}carbonyl]phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.02 (m, 1H), 8.99 (m, 1H), 8.93 (m, 1H), 8.75 (m, 2H), 8.41 (m, 1H), 8.31 (m, 1H), 7.78 (m, 1H), 7.54 (m, 1H), 7.50 (m, 1H), 7.35 (m, 4H), 7.25 (m, 4H), 7.15 (m, 2H), 5.32 (m, 2H), 2.67 (m, 2H), 2.46 (m, 2H), 2.33 (m, 2H), 1.92 (m, 2H), 1.73 (m, 2H). MS (EI) for $C_{31}H_{32}N_6O_2$: 521 (MH^+).

[1186] 3-Amino-6-[3-({[(3,5-difluorophenyl)methyl]amino}carbonyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.43 (m, 1H), 8.96 (s, 2H), 8.85 (m, 1H), 8.64 (s, 1H), 8.35 (m, 1H), 7.92 (m, 1H), 7.61 (m, 1H), 7.10 (m, 3H), 4.53 (m, 2H), 3.68 (m, 2H), 3.60 (m, 1H), 3.48 (m, 2H), 3.25 (m, 2H), 1.91 (m, 2H), 1.77 (m, 2H). MS (EI) for $C_{24}H_{24}F_2N_6O_2$: 467 (MH^+).

[1187] Methyl 4-[{[3-(5-amino-6-{[(3*S*)-piperidin-3-ylamino]carbonyl}pyrazin-2-yl)phenyl]carbonyl}amino]methyl]benzoate: 1H NMR (400 MHz, d_6 -DMSO): δ 9.39 (m, 1H), 8.96 (s, 1H), 8.81 (d, $J = 8.8$ Hz, 1H), 8.36 (d, $J = 8.0$ Hz, 1H), 7.95 (d, $J = 8.4$ Hz, 2H), 7.92 (d, $J = 7.6$ Hz, 1H), 7.61 (t, $J = 7.6$ Hz, 1H), 7.52 (d, $J = 8.4$ Hz, 2H), 4.61 (d, $J = 6.0$ Hz, 2H), 3.85 (s, 3H), 3.67 (m, 1H), 3.48 (m, 2H), 3.25 (m, 2H), 3.08 (m, 2H), 2.77 (m, 2H), 1.90 (m, 2H), 1.75 (m, 2H). MS (EI) for $C_{26}H_{28}N_6O_4$: 489 (MH^+).

- [1188] 3-Amino-6-[3-(((4-bromophenyl)methyl)amino)carbonyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.47 (m, 1H), 9.11 (m, 1H), 8.95 (s, 1H), 8.89 (d, $J = 8.8$ Hz, 1H), 8.67 (s, 1H), 8.34 (d, 8.4 Hz, 1H), 7.90 (d, $J = 8.0$ Hz, 1H), 7.59 (t, $J = 7.6$ Hz, 1H), 7.52 (m, 2H), 7.34 (d, $J = 8.4$ Hz, 2H), 4.49 (d, $J = 6.0$ Hz, 2H), 3.69 (m, 2H), 3.48 (m, 2H), 3.24 (m, 2H), 2.77 (m, 1H), 1.86 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{25}\text{N}_6\text{O}_2\text{Br}$: 510 (MH^+).
- [1189] 3-Amino-6-[3-(((4-bromo-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.45 (m, 1H), 9.14 (m, 1H), 8.95 (s, 1H), 8.67 (s, 1H), 8.35 (d, $J = 8.0$ Hz, 1H), 7.90 (d, $J = 7.6$ Hz, 1H), 7.58 (m, 2H), 7.41 (m, 2H), 4.51 (d, $J = 8.0$ Hz, 2H), 4.28 (m, 1H), 3.60 (m, 1H), 3.24 (m, 2H), 2.77 (m, 2H), 1.89 (m, 2H), 1.81 (m, 2H). MS (EI) for $\text{C}_{24}\text{H}_{24}\text{BrFN}_6\text{O}_2$: 528 (MH^+).
- [1190] 3-Amino-6-[3-(((2-bromophenyl)methyl)amino)carbonyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.39 (t, $J = 5.6$ Hz, 1H), 9.00 (m, 1H), 8.97 (s, 1H), 8.85 (m, 1H), 8.68 (s, 1H), 8.36 (m, 1H), 7.93 (m, 1H), 7.64 (m, 2H), 7.39 (m, 2H), 7.23 (m, 1H), 4.51 (d, $J = 6.0$ Hz, 2H), 3.67 (m, 1H), 3.50 (m, 1H), 3.25 (m, 2H), 3.14 (m, 1H), 2.77 (m, 2H), 1.91 (m, 2H), 1.76 (m, 2H). MS (EI) for $\text{C}_{24}\text{H}_{25}\text{BrN}_6\text{O}_2$: 510 (MH^+).
- [1191] 3-Amino-6-[3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.2 (t, 1H), 8.9 (s, 1H), 8.55 (s, 1H), 8.45 (d, 1H), 8.3 (d, 1H), 7.9 (d, 1H), 7.7-7.5 (m, 5H), 4.7 (d, 2H), 3.85 (m, 1H), 2.9 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_6\text{O}_2\text{F}_4$: 517 (MH^+).
- [1192] 3-Amino-6-[3-(((2-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.3 (t, 1H), 8.95 (s, 1H), 8.6 (d, 1H), 8.55 (s, 1H), 8.3 (d, 1H), 8.0-7.5 (m, 6H), 7.4 (t, 1H), 4.6 (d, 2H), 4.0 (m, 1H), 3.1-2.9 (m, 2H), 2.75 (t, 1H), 2.6 (t, 1H), 1.9-1.5 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_6\text{O}_2\text{F}_4$: 517 (MH^+).
- [1193] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-[3-(((3*R*)-pyrrolidin-3-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.83 (s, 1H), 8.72 (s, 1H), 8.43 (m, 1H), 8.17 (m, 1H), 7.79 (m, 1H), 7.51 (m, 1H), 4.26 (m, 1H),

3.65 (m, 1H), 3.58 (m, 2H), 3.47 (m, 3H), 3.37 (m, 3H), 3.12 (m, 1H), 2.99 (m, 2H), 2.68 (m, 1H), 2.13 (m, 1H), 2.00 (m, 2H), 1.83 (m, 3H). MS (EI) for $C_{22}H_{29}N_7O_2$: 424 (MH^+).

[1194] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([(3*R*)-pyrrolidin-3-ylamino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.80 (s, 1H), 8.58 (s, 1H), 8.24 (d, $J = 8.0$ Hz, 1H), 7.91 (d, $J = 8.4$ Hz, 1H), 7.60 (t, $J = 8.0$ Hz, 1H), 4.33 (m, 1H), 3.75 (m, 1H), 3.63 (m, 2H), 3.50 (m, 2H), 3.20 (m, 2H), 3.04 (m, 2H), 2.43 (m, 2H), 2.28 (m, 2H), 2.10 (m, 2H), 1.92 (m, 2H). MS (EI) for $C_{21}H_{27}N_7O_2$: 410 (MH^+).

[1195] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([(2-piperidin-1-ylethyl)amino]carbonyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.80 (s, 1H), 8.64 (s, 1H), 8.24 (d, $J = 7.6$ Hz, 1H), 7.94 (d, $J = 7.6$ Hz, 1H), 7.61 (t, $J = 7.6$ Hz, 1H), 4.33 (m, 1H), 3.85 (m, 2H), 3.69 (m, 5H), 3.58 (m, 1H), 3.47 (m, 1H), 3.39 (m, 2H), 3.24 (m, 2H), 3.02 (m, 2H), 2.10 (m, 2H), 2.00 (m, 2H), 1.86 (m, 4H). MS (EI) for $C_{24}H_{33}N_7O_2$: 452 (MH^+).

[1196] 3-Amino-6-(3-([(2-phenylethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.76 (s, 1H), 8.48 (s, 1H), 8.18 (d, $J = 7.6$ Hz, 1H), 7.78 (d, $J = 7.6$ Hz, 1H), 7.76 (t, $J = 7.6$ Hz, 1H), 7.26 (m, 4H), 7.19 (m, 1H), 4.34 (m, 1H), 3.73 (m, 1H), 3.66 (m, 4H), 3.57 (m, 1H), 3.48 (m, 1H), 3.36 (m, 2H), 3.15 (m, 1H), 3.02 (m, 1H), 2.95 (m, 2H), 2.09 (m, 2H), 1.90 (m, 2H). MS (EI) for $C_{25}H_{28}N_6O_2$: 445 (MH^+).

[1197] 3-Amino-6-[3-([(2-(methyloxy)ethyl)amino]carbonyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.76 (s, 1H), 8.48 (s, 1H), 8.18 (d, $J = 7.6$ Hz, 1H), 7.78 (d, $J = 7.6$ Hz, 1H), 7.76 (t, $J = 7.6$ Hz, 1H), 4.33 (m, 1H), 3.74 (m, 1H), 3.67 (m, 2H), 3.59 (m, 1H), 3.49 (m, 1H), 3.39 (m, 3H), 3.34 (m, 2H), 3.16 (m, 1H), 3.02 (m, 1H), 2.10 (m, 2H), 1.90 (m, 2H). MS (EI) for $C_{20}H_{26}N_6O_3$: 399 (MH^+).

[1198] 3-Amino-6-[3-([(4-iodophenyl)methyl]amino]carbonyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 8.74 (s, 1H), 8.58 (m, 1H), 8.24-8.21 (m, 1H), 7.90-7.88 (m, 1H), 7.62-7.59 (m, 1H), 7.39-7.32 (m, 4H), 4.60 (s, 2H), 4.31-4.29 (m, 1H), 3.50-3.46 (m, 2H), 3.37-3.30 (m, 2H), 2.10-1.83 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_2I$: 557 (MH^+).

- [1199] 3-Amino-6-[3-({[(2-chlorophenyl)methyl]amino}carbonyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 8.78 (s, 1H), 8.57 (m, 1H), 8.23-8.21 (m, 1H), 7.92-7.89 (m, 1H), 7.62-7.59 (m, 1H), 7.37-7.30 (m, 4H), 4.60 (s, 2H), 4.31-4.28 (m, 1H), 3.50-3.46 (m, 2H), 3.37-3.30 (m, 2H), 2.10-1.83 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{25}\text{N}_6\text{O}_2\text{Cl}$: 465 (MH^+).
- [1200] 3-Amino-6-(3-({[(4-chlorophenyl)acetyl]amino}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.8 (s, 1H), 9.2 (m, 1H), 9.03 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.42 (s, 1H), 7.8 (m, 2H), 7.4 (m, 5H), 4.3 (m, 1H), 3.7 (s, 2H), 3.3-3.2 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{25}\text{N}_6\text{O}_2\text{Cl}$: 465 (MH^+).
- [1201] 3-Amino-6-(3-({[(2,6-dichlorophenyl)acetyl]amino}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.8 (s, 1H), 9.2 (m, 1H), 9.0 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.4 (s, 1H), 7.85 (d, 1H), 7.72 (d, 1H), 7.5 (t, 1H), 7.4 (m, 3H), 4.3 (m, 1H), 4.1 (s, 2H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2\text{Cl}_2$: 499 (MH^+).
- [1202] 3-Amino-6-(3-({[(pentafluorophenyl)acetyl]amino}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.95 (s, 1H), 9.2 (m, 1H), 9.05 (m, 1H), 8.78 (s, 1H), 8.65 (d, 1H), 8.4 (s, 1H), 7.85 (d, 1H), 7.75 (d, 1H), 7.4 (t, 1H), 4.3 (m, 1H), 3.96 (s, 2H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{21}\text{N}_6\text{O}_2\text{F}_5$: 521 (MH^+).
- [1203] 3-Amino-6-(3-({[(2-chloro-4-fluorophenyl)acetyl]amino}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.2 (m, 1H), 9.05 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.4 (s, 1H), 7.85 (d, 1H), 7.75 (d, 1H), 7.5 (t, 1H), 7.45 (m, 2H), 7.25 (m, 1H), 4.3 (m, 1H), 3.9 (s, 2H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2\text{FCl}$: 483 (MH^+).
- [1204] 3-Amino-6-[3-({[4-(methylthio)phenyl]acetyl]amino}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.2 (m, 1H), 9.0 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.4 (s, 1H), 7.8 (d, 2H), 7.4 (m, 3H), 7.2 (d, 2H), 4.3 (m, 1H), 3.7 (s, 2H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.45 (s, 3H), 2.0-1.7 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2\text{S}$: 477 (MH^+).
- [1205] 3-Amino-6-(3-({[(2,4-dichlorophenyl)acetyl]amino}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 10.8 (s, 1H), 9.25 (m,

1H), 9.1 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.43 (s, 1H), 7.85 (d, 1H), 7.8 (d, 1H), 7.6 (s, 1H), 7.5 (t, 1H), 7.4 (m, 2H), 4.3 (m, 1H), 3.9 (s, 2H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{24}H_{24}N_6O_2Cl_2$: 499 (MH^+).

[1206] 3-Amino-6-[3-({[4-(4-chlorophenyl)-2-thienyl]carbonyl}amino)phenyl]-*N*-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.85 (s, 1H), 9.2-9.1 (m, 2H), 9.0-8.8 (m, 4H), 8.27 (s, 1H), 8.05 (d, 1H), 7.9-7.8 (m, 3H), 7.57 (d, 2H), 7.45 (t, 1H), 4.4 (m, 1H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{27}H_{25}N_6O_2SCl$: 533 (MH^+).

[1207] 3-Amino-6-(3-({[2,4-difluorophenyl]acetyl}amino)phenyl)-*N*-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.1 (m, 1H), 8.95 (m, 1H), 8.75 (s, 1H), 8.67 (d, 1H), 8.4 (s, 1H), 7.82 (d, 1H), 7.72 (d, 1H), 7.5 (q, 1H), 7.4 (t, 1H), 7.25 (m, 1), 7.1 (m, 1H), 4.5 (s, 1H), 4.3 (m, 1H), 3.8 (s, 2H), 3.2 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{24}H_{24}N_6O_2F_2$: 483 (MH^+).

[1208] 3-Amino-6-(3-({[2-fluorophenyl]acetyl}amino)phenyl)-*N*-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.15 (m, 1H), 9.0 (m, 1H), 8.75 (s, 1H), 8.7 (d, 1H), 8.4 (s, 1H), 7.8 (d, 1H), 7.75 (d, 1H), 7.5-7.1 (m, 5H), 4.3 (m, 1H), 3.8 (s, 2H), 3.2 (m, 3H), 2.75 (s, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_2F$: 449 (MH^+).

[1209] 3-Amino-6-[3-({[5-[2-chloro-5-(trifluoromethyl)phenyl]furan-2-yl]carbonyl}amino)phenyl]-*N*-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.75 (s, 1H), 9.05 (m, 1H), 8.9 (m, 1H), 8.85 (s, 1H), 8.75 (d, 1H), 8.6 (s, 1H), 8.5 (s, 1H), 7.9 (t, 3H), 7.8 (m, 2H), 7.5 (m, 2H), 4.3 (m, 1H), 3.35 (m, 3H), 2.75 (q, 1H), 1.9 (m, 3H), 1.75 (m, 1H). MS (EI) for $C_{28}H_{24}N_6O_3F_3Cl$: 585 (MH^+).

[1210] 3-Amino-6-[3-({[4-(dimethylamino)naphthalen-1-yl]carbonyl}amino)phenyl]-*N*-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.2 (m, 1H), 9.05 (m, 1H), 8.8 (s, 1H), 8.7 (d, 1H), 8.55 (s, 1H), 8.3 (m, 2H), 7.9 (t, 2H), 7.8 (d, 1H), 7.6 (m, 2H), 7.5 (t, 1H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 3.0 (s, 6H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{29}H_{31}N_7O_2$: 510 (MH^+).

[1211] 3-Amino-*N*-[(3S)-piperidin-3-yl]-6-[3-({[4-(1,2,3-thiadiazol-5-yl)phenyl]carbonyl}amino)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.85 (s, 1H), 9.15 (m, 1H), 8.95 (m, 1H), 8.8 (d, 2H), 8.67 (s,

1H), 8.3 (q, 4H), 8.0 (d, 1H), 7.9 (d, 1H), 7.5 (t, 1H), 4.3 (m, 1H), 3.25 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{25}H_{24}N_8O_2S$: 501 (MH^+).

[1212] 3-Amino-6-(3-([(1,3-dimethyl-1H-pyrazol-5-yl)carbonyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.5 (s, 1H), 9.25 (m, 1H), 9.1 (m, 1H), 8.8 (m, 2H), 8.7 (s, 1H), 7.95 (d, 1H), 7.85 (d, 1H), 7.45 (t, 1H), 7.2 (s, 1H), 5.75 (s, 1H), 4.3 (m, 1H), 3.25 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{22}H_{26}N_8O_2$: 435 (MH^+).

[1213] 3-Amino-6-(3-([(4-bromo-2-fluorophenyl)carbonyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.7 (s, 1H), 9.1 (m, 1H), 8.9 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.4 (s, 1H), 7.9 (d, 1H), 7.8 (m, 2H), 7.7 (d, 1H), 7.58 (d, 1H), 7.5 (t, 1H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{23}H_{22}N_6O_2FBr$: 515 (MH^+).

[1214] N-[3-(5-Amino-6-([(3S)-piperidin-3-ylamino]carbonyl)pyrazin-2-yl)phenyl]quinoline-3-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 11.15 (s, 1H), 9.65 (s, 1H), 9.4 (s, 1H), 9.18 (m, 1H), 8.98 (m, 1H), 8.83 (s, 1H), 8.8 (m, 2H), 8.25 (d, 1H), 8.2 (d, 1H), 8.1 (d, 1H), 8.0 (t, 1H), 7.9 (d, 1H), 7.8 (t, 1H), 7.5 (t, 1H), 4.3 (m, 1H), 3.3-3.2 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{26}H_{25}N_7O_2$: 468 (MH^+).

[1215] 3-Amino-6-[3-([(6-(methyloxy)-1-benzofuran-3-yl)acetyl]amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.6 (s, 1H), 8.9 (m, 1H), 8.8 (m, 1H), 8.78 (s, 1H), 8.7 (d, 1H), 8.35 (s, 1H), 7.8 (m, 2H), 7.73 (d, 1H), 7.6 (d, 1H), 7.4 (t, 1H), 7.2 (s, 1H), 6.9 (d, 1H), 4.3 (m, 1H), 3.8 (s, 2H), 3.78 (d, 2H), 3.3-3.1 (m, 3H), 2.8 (m, 1H), 2.0-1.7 (m, 4H). MS (EI) for $C_{27}H_{28}N_6O_4$: 501 (MH^+).

[1216] 3-Amino-N-[(3S)-piperidin-3-yl]-6-[3-([(4'-(trifluoromethyl)biphenyl-2-yl)carbonyl]amino)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.5 (s, 1H), 9.05 (m, 1H), 8.9 (m, 1H), 8.67 (s, 1H), 8.6 (d, 1H), 8.2 (s, 1H), 7.85 (d, 1H), 7.75 (d, 2H), 7.7-7.5 (m, 7H), 7.4 (t, 1H), 4.3 (m, 1H), 4.05 (s, 1H), 3.25 (m, 2H), 3.05 (m, 1H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{30}H_{27}N_6O_2F_3$: 561 (MH^+).

[1217] 3-Amino-6-[3-([(3,5-bis(trifluoromethyl)phenyl)acetyl]amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 0.8 (s, 1H), 9.05 (m, 1H), 8.9 (m, 1H), 8.72 (s, 1H), 8.52 (d, 1H), 8.35 (s, 1H), 8.1 (s, 2H), 8.0 (s, 1H), 7.8 (d, 1H), 7.7 (d, 2H), 7.4 (t, 1H), 4.3 (m, 1H), 4.0 (s, 2H), 3.3-3.15 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for $C_{26}H_{24}N_6O_2F_6$: 567 (MH^+).

- [1218] 3-Amino-6-{3-[[[4-(4-methylpiperazin-1-yl)phenyl]methyl]amino]carbonyl}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 8.60 (s, 1H), 8.23 (m, 1H), 7.90 (m, 1H), 7.61 (m, 1H), 7.36 (m, 2H), 7.06 (m, 2H), 4.56 (s, 2H), 4.39-4.37 (m, 1H), 3.86-3.83 (m, 2H), 3.67-3.61 (m, 2H), 3.51-3.48 (m, 2H), 3.38-3.35 (m, 1H), 3.31-2.97 (m, 8H), 2.09-1.96 (m, 4H). MS (EI) for $\text{C}_{29}\text{H}_{36}\text{N}_8\text{O}_2$: 529 (MH^+).
- [1219] 3-Amino-6-[3-([2-(4-fluorophenyl)ethyl]amino)carbonyl]phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.74 (s, 1H), 8.50 (m, 1H), 8.21-8.19 (m, 1H), 7.81-7.78 (m, 1H), 7.59-7.55 (m, 1H), 7.30-7.26 (m, 2H), 7.03-6.99 (m, 2H), 4.35 (m, 1H), 3.69-2.93 (m, 8H), 2.11-1.91 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{27}\text{N}_6\text{O}_2\text{F}$: 463 (MH^+).
- [1220] 3-Amino-6-(3-[(2-morpholin-4-ylethyl)amino]carbonyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.80 (s, 1H), 8.67 (s, 1H), 8.24 (m, 1H), 7.95 (m, 1H), 7.63-7.59 (m, 1H), 4.37-4.36 (m, 1H), 4.12-4.08 (m, 2H), 3.89-3.84 (m, 4H), 3.75-3.66 (m, 2H), 3.49-3.03 (m, 8H), 2.12-1.91 (m, 4H). MS (EI) for $\text{C}_{23}\text{H}_{31}\text{N}_7\text{O}_3$: 454 (MH^+).
- [1221] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-[3-([2*R*)-pyrrolidin-2-ylmethyl]amino]carbonyl}phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.77 (s, 1H), 8.61 (s, 1H), 8.25-8.23 (m, 1H), 7.93-7.91 (m, 1H), 7.63-7.61 (m, 1H), 4.37 (m, 1H), 3.57-2.77 (m, 11H), 2.24-1.86 (m, 6H). MS (EI) for $\text{C}_{22}\text{H}_{29}\text{N}_7\text{O}_2$: 424 (MH^+).
- [1222] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-[(2*S*)-pyrrolidin-2-ylamino]carbonyl}phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.77 (s, 1H), 8.61 (s, 1H), 8.25-8.23 (m, 1H), 7.92-7.89 (m, 1H), 7.64-7.62 (m, 1H), 4.78-4.75 (m, 1H), 4.39-4.37 (m, 1H), 3.57-2.77 (m, 8H), 2.44-1.93 (m, 6H). MS (EI) for $\text{C}_{21}\text{H}_{27}\text{N}_7\text{O}_2$: 410 (MH^+).
- [1223] 3-Amino-6-{3-[(cyclopentylamino)carbonyl]phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.73 (s, 1H), 8.53 (s, 1H), 8.24-8.22 (m, 1H), 7.88-7.86 (m, 1H), 7.62-7.58 (m, 1H), 4.99-4.89 (m, 2H), 3.59-3.49 (m, 1H), 3.39-3.30 (m, 1H), 3.24-3.21 (m, 1H), 3.09-3.01 (m, 1H), 2.37-1.67 (m, 11H). MS (EI) for $\text{C}_{22}\text{H}_{27}\text{N}_6\text{O}_2$: 409 (MH^+).

- [1224] 3-Amino-6-{3-([4-(1,1-dimethylethyl)phenyl]methyl)amino}carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.89 (s, 1H), 8.57 (m, 1H), 8.18 (m, 1H), 8.05 (m, 1H), 7.82 (m, 1H), 7.78 (m, 1H), 7.42-7.24 (m, 4H), 4.62 (m, 3H), 4.27 (m, 1H), 3.42-2.92 (m, 4H), 2.18-1.84 (m, 4H), 1.27 (s, 9H). MS (EI) for C₂₈H₃₄N₆O₂: 487 (MH⁺).
- [1225] 3-Amino-6-[3-([(2,4-dichloro-6-methylphenyl)methyl]amino)carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.78 (s, 1H), 8.65 (m, 1H), 8.18 (m, 1H), 7.82 (m, 1H), 7.57 (m, 1H), 7.39-7.23 (m, 2H), 4.68 (m, 3H), 4.26 (m, 1H), 3.42-2.92 (m, 4H), 2.44 (s, 3H), 2.19-1.86 (m, 4H). MS (EI) for C₂₅H₂₆N₆O₂Cl₂: 513 (MH⁺).
- [1226] 3-Amino-6-[3-([(3,5-dichlorophenyl)methyl]amino)carbonylphenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.82 (s, 1H), 8.63 (m, 1H), 8.57 (m, 1H), 8.22 (m, 1H), 7.59 (m, 1H), 7.26 (s, 3H), 4.68 (m, 3H), 4.26 (m, 1H), 3.47-2.99 (m, 4H), 2.15-1.82 (m, 4H). MS (EI) for C₂₄H₂₄N₆O₂Cl₂: 499 (MH⁺).
- [1227] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-{3-([4-(1,2,3-thiadiazol-4-yl)phenyl]methyl)amino}carbonylphenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.6 (s, 1H), 9.55 (t, 1H), 9.2 (m, 1H), 9.0-8.9 (m, 3H), 8.7 (s, 1H), 8.35 (d, 1H), 8.1 (d, 2H), 7.95 (d, 1H), 7.6-7.5 (m, 3H), 4.6 (d, 2H), 4.3 (m, 1H), 3.3-3.1 (m, 3H), 2.75 (m, 1H), 1.95-1.7 (m, 4H). MS (EI) for C₂₆H₂₆N₈O₂S: 515 (MH⁺).
- [1228] 3-Amino-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 10.0 (s, 2H), 9.55 (s, 2H), 9.15 (d, 1H), 8.92 (s, 1H), 8.12 (d, 1H), 7.9 (d, 1H), 7.6-7.45 (m, 4H), 7.35-7.25 (m, 3H), 4.85-4.8 (m, 2H), 4.3 (s, 2H), 3.4 (m, 4H), 3.2 (m, 2H), 2.9 (m, 1H), 2.45 (m, 2H), 2.3-2.15 (m, 2H). MS (EI) for C₂₅H₂₈N₆O: 429 (MH⁺).
- [1229] 3-Amino-6-[3-([(1*R*,2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino)methyl]phenyl]-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 10.2 (s, 1H), 9.6 (s, 2H), 9.3 (m, 1H), 9.1 (d, 1H), 8.9 (d, 2H), 8.15 (d, 1H), 7.9 (d, 1H), 7.58 (d, 1H), 7.45 (t, 1H), 7.3-7.2 (m, 3H), 4.8 (m, 2H), 4.6 (m, 1H), 4.45 (m, 2H), 3.55 (m, 3H), 3.25-3.1 (m, 2H), 3.0 (d, 1H), 2.3-2.15 (m, 2H). MS (EI) for C₂₅H₂₈N₆O₂: 445. (MH⁺).
- [1230] 3-Amino-*N*-[(3*S*)-pyrrolidin-3-yl]-6-(3-([(1*S*)-1,2,3,4-tetrahydronaphthalen-1-ylamino]methyl)phenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, CDCl₃): δ 8.63

(s, 1H), 8.24 (d, 1H), 7.75 (s, 1H), 7.72-7.75 (m, 1H), 7.38-7.46 (m, 3H), 7.14-7.18 (m, 2H), 7.08-7.11 (m, 1H), 4.52-4.56 (m, 1H), 3.94-4.06 (m, 2H), 3.09 (t, 1H), 2.96-3.03 (m, 2H), 2.68-2.88 (m, 5H), 2.21-2.29 (m, 1H), 1.94-2.06 (m, 3H), 1.74-1.86 (m, 2H). MS (EI) for $C_{26}H_{30}N_6O$: 443 (MH^+).

[1231] 3-Amino-6-(3-hydroxyphenyl)-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide:

1H NMR (400 MHz, d_6 -DMSO): δ 9.36 (br s, 2H), 8.80 (d, 1H), 8.75 (s, 1H), 7.57-7.53 (m, 2H), 7.23 (t, 1H), 6.78 (d, 1H), 4.64-4.57 (m, 2H), 4.20 (br s, 1H), 3.44-3.34 (m, 2H), 3.29-3.17 (m, 2H), 2.28-2.19 (m, 1H), 2.10-2.03 (m, 1H). MS (EI) for $C_{15}H_{17}N_7O_2$: 300 (MH^+).

[1232] 3-Amino-6-(3-hydroxyphenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide:

1H NMR (400 MHz, d_6 -DMSO): δ 9.54 (br s, 1H), 8.93 (m, 1H), 8.82-8.79 (m, 1H), 7.75 (s, 1H), 8.62 (d, 1H), 7.60 (br s, 1H), 7.59-7.46 (m, 2H), 7.24 (t, 1H), 6.81 (d, 1H), 4.21 (br s, 1H), 3.27-3.20 (m, 2H), 3.03 (q, 1H), 2.76-2.74 (m, 1H), 1.89-1.87 (m, 2H), 1.76-1.72 (m, 2H). MS (EI) for $C_{16}H_{19}N_5O_2$: 314 (MH^+).

[1233] 5-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]methyl)phenyl)-3-(4-piperidin-3-yl-1,3-thiazol-2-yl)pyrazin-2-amine:

1H NMR (400 MHz, d_6 -DMSO): δ 9.4 (t, 2H), 8.9 (d, 1H), 8.8 (s, 1H), 8.63 (d, 1H), 8.15 (s, 1H), 8.05 (d, 1H), 7.9 (m, 1H), 7.67 (m, 2H), 7.6-7.5 (m, 2H), 7.4-7.25 (m, 3H), 4.85 (m, 1H), 4.3 (d, 2H), 3.6 (d, 1H), 3.4-3.0 (m, 4H), 2.9 (t, 2H), 2.4-2.0 (m, 2H), 2.0-1.7 (m, 3H). MS (EI) for $C_{28}H_{30}N_6S$: 483 (MH^+).

[1234] 1,1-Dimethylethyl (3*S*)-3-([(3-amino-6-{3-([(1,1-dimethylethyl)oxy]carbonyl)amino)methyl}phenyl]pyrazin-2-yl)carbonyl]amino)piperidine-1-carboxylate:

1H NMR (400 MHz, $CDCl_3$): δ 8.4 (s, 1H), 8.2 (m, 1H), 7.80 (m, 2H), 7.4 (m, 1H), 4.6 (s, 2H), 4.1 (m, 1H), 3.4 (m, 4H), 1.8 (m, 4H), 1.5-1.6 (m, 18H). MS (EI) for $C_{27}H_{38}N_6O_5$: 527 (MH^+).

[1235] 3-Amino-6-[3-(aminomethyl)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-

carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.4 (s, 1H), 8.2 (m, 1H), 7.80 (m, 2H), 7.4 (m, 1H), 4.6 (s, 2H), 4.1 (m, 1H), 3.4 (m, 4H), 1.8 (m, 4H). MS (EI) for $C_{17}H_{22}N_6O$: 327 (MH^+).

[1236] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-{3-([(2-(2-thienyl)-1,3-thiazol-4-

yl)carbonyl]amino)methyl}phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.4 (s, 1H), 8.2 (m, 1H), 7.80 (m, 2H), 7.6 (s, 1H), 7.4 (m, 3H), 7.2 (m, 1H)

4.6 (s, 2H), 4.1 (m, 1H), 3.4 (m, 4H), 1.8 (m, 4H). MS (EI) for $C_{25}H_{25}N_7O_2S_2$: 520 (MH^+).

[1237] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-{3-[[[5-(2-thienyl)pyridin-3-yl]carbonyl]amino)methyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.4 (s, 1H), 8.2 (m, 1H), 7.80 (m, 2H), 7.6 (s, 1H), 7.4 (m, 3H), 7.3 (m, 2H), 7.2 (m, 1H), 4.6 (s, 2H), 4.1 (m, 1H), 3.4 (m, 4H), 1.8 (m, 4H). MS (EI) for $C_{27}H_{27}N_7O_2S$: 514 (MH^+).

[1238] 1,1-Dimethylethyl (3*S*)-3-[[[3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate: 1H NMR (400 MHz, d_6 -DMSO): δ 8.95 (m, 1H), 8.50 (d, 1H), 8.40-8.10 (m, 4H), 7.60-7.40 (m, 2H), 4.10 (m, 2H), 3.90-3.60 (m, 3H), 3.30-3.00 (m, 2H), 1.90-1.60 (m, 4H), 1.40 (s, 9H). MS (EI) for $C_{22}H_{30}N_6O_3$: 427 (MH^+).

[1239] 3-Amino-6-[3-[[[4-fluorophenyl]carbonyl]amino]methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 8.02 (s, 1H), 7.90-7.95 (m, 3H), 7.40-7.46 (m, 2H), 7.21 (t, 2H), 4.66 (m, 2H), 3.37-3.73 (m, 2H), 2.91-3.13 (m, 3H), 1.76-2.06 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_2F$: 449 (MH^+).

[1240] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-[3-[[[4-[(trifluoromethyl)oxy]phenyl]carbonyl]amino]methyl]phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 9.14-9.20 (m, 1H), 8.72 (s, 1H), 8.58-8.59 (m, 1H), 8.03 (s, 1H), 7.97-7.99 (m, 2H), 7.91-7.93 (m, 1H), 7.41-7.48 (m, 1H), 7.38-7.39 (m, 1H), 4.62-4.71 (m, 2H), 4.27 (s, 1H), 3.47-3.50 (m, 1H), 3.32-3.49 (m, 1H), 3.06-3.12 (m, 2H), 2.94-2.99 (m, 1H), 2.03-2.06 (m, 2H), 1.79-1.87 (m, 2H). MS (EI) for $C_{25}H_{25}N_6O_3F_3$: 515 (MH^+).

[1241] 3-Amino-6-[3-[[[4-chlorophenyl]carbonyl]amino]methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 8.02 (s, 1H), 7.90-7.92 (m, 2H), 7.85-7.87 (m, 2H), 7.44-7.51 (m, 3H), 7.38-7.39 (m, 1H), 4.64-4.66 (m, 2H), 4.27 (s, 1H), 3.46-3.73 (m, 2H), 2.96-3.09 (m, 3H), 1.77-2.05 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_2Cl$: 465 (MH^+).

[1242] 3-Amino-6-[3-[[[4-bromophenyl]carbonyl]amino]methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 8.01 (s, 1H), 7.93-7.89 (m, 1H), 7.81-7.76 (m, 2H), 7.68-7.63 (m, 2H), 7.48-7.42 (m, 1H), 7.41-7.36 (m, 1H), 4.71-4.59 (m, 2H), 4.35-4.20 (m, 1H), 3.51-3.44 (m, 1H), 3.39-3.30

(m, 1H), 3.11-3.02 (m, 1H), 3.00-2.91 (m, 1H), 2.10-1.96 (m, 2H), 1.94-1.67 (m, 2H). MS (EI) for $C_{24}H_{25}N_6O_2Br$: 508 (MH^+).

- [1243] 3-Amino-6-(3-([(naphthalen-2-ylcarbonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.73 (s, 1H), 8.45 (s, 1H), 8.07 (s, 1H), 8.02-7.88 (m, 5H), 7.64-7.54 (m, 2H), 7.51-7.40 (m, 2H), 4.81-4.65 (m, 2H), 4.25-4.12 (m, 1H), 3.49-3.40 (m, 1H), 3.30-3.22 (m, 1H), 3.02-2.93 (m, 1H), 2.84-2.74 (m, 1H), 1.95-1.65 (m, 3H), 1.63-1.50 (m, 1H). MS (EI) for $C_{28}H_{28}N_6O_2$: 481 (MH^+).
- [1244] 3-Amino-6-{3-([(4-(dimethylamino)phenyl)carbonyl]amino)methyl}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.55 (d, 1H), 8.00 (s, 1H), 7.90-7.70 (m, 3H), 7.50-7.30 (m, 2H), 6.80-6.70 (m, 2H), 4.65 (m, 2H), 3.50-2.80 (m, 10H), 2.00-1.60 (m, 4H). MS (EI) for $C_{26}H_{31}N_7O_2$: 474 (MH^+).
- [1245] 3-Amino-6-{3-([(4-(methyloxy)phenyl)carbonyl]amino)methyl}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.55 (d, 1H), 8.05 (s, 1H), 8.00-7.85 (m, 4H), 7.50-7.40 (m, 2H), 7.00 (m, 2H), 4.65 (m, 2H), 4.25 (m, 1H), 3.80 (s, 3H), 3.50 (m, 1H), 3.10 (m, 1H), 2.95 (m, 1H), 2.00 (m, 2H), 1.80 (m, 2H). MS (EI) for $C_{25}H_{28}N_6O_3$: 461 (MH^+).
- [1246] 3-Amino-6-(3-([(biphenyl-4-ylcarbonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.55 (d, 1H), 8.05 (s, 1H), 8.00-7.85 (m, 3H), 7.80-7.60 (m, 4H), 7.50-7.30 (m, 5H), 4.65 (m, 2H), 4.25 (m, 1H), 3.50 (m, 1H), 3.10 (m, 1H), 2.95 (m, 1H), 2.00 (m, 2H), 1.80 (m, 2H). MS (EI) for $C_{30}H_{30}N_6O_2$: 507 (MH^+).
- [1247] 3-Amino-6-(3-([(furan-2-ylcarbonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.55 (d, 1H), 8.10 (s, 1H), 8.00 (s, 1H), 7.95 (m, 1H), 7.60 (m, 1H), 7.45-7.25 (m, 2H), 6.80 (m, 1H), 4.65 (m, 2H), 4.25 (m, 1H), 3.50 (m, 1H), 2.95 (m, 3H), 2.00 (m, 2H), 1.80 (m, 2H). MS (EI) for $C_{22}H_{24}N_6O_3$: 421 (MH^+).
- [1248] 3-Amino-6-(3-([(2,3-dihydro-1-benzofuran-5-ylcarbonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 8.01 (s, 1H), 7.92-7.87 (m, 1H), 7.76 (s, 1H), 7.71-7.66 (m, 1H), 7.48-7.41 (m, 1H), 7.40-7.35 (m, 1H), 6.81-6.76 (m, 1H), 4.67-4.57 (m, 4H), 4.33-4.20 (m, 1H), 3.52-

- 3.42 (m, 1H), 3.40-3.20 (m, 3H), 3.11-3.02 (m, 1H), 3.00-2.90 (m, 1H), 2.08-1.97 (m, 2H), 1.90-1.68 (m, 2H). MS (EI) for $C_{26}H_{28}N_6O_3$: 473 (MH^+).
- [1249] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([(2-thienylcarbonyl)amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.55, d, 1H), 8.10 (s, 1H), 8.00 (s, 1H), 7.95 (m, 1H), 7.60, 7.35 (m, 5H), 4.65 (m, 2H), 4.25 (m, 1H), 3.50 (m, 1H), 2.95 (m, 3H), 2.00 (m, 2H), 1.80 (m, 2H). MS (EI) for $C_{22}H_{24}N_6O_2S$: 437 (MH^+).
- [1250] 3-Amino-6-(3-([(2,3-dihydro-1,4-benzodioxin-2-ylcarbonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.80 (m, 1H), 8.70 (m, 1H), 7.95-7.75 (m, 2H), 7.40 (m, 1H), 7.20 (m, 1H), 7.00 (m, 1H), 6.95-6.85 (m, 3H), 4.85 (m, 1H), 4.60 (m, 1H), 4.50-4.25 (m, 4H), 3.50 (m, 1H), 3.35 (m, 1H), 3.10 (m, 1H), 2.95 (m, 1H), 2.10 (m, 2H), 1.95 (m, 2H). MS (EI) for $C_{26}H_{28}N_6O_4$: 489 (MH^+).
- [1251] *N*-[(3-{5-Amino-6-[(3*R*)-piperidin-3-ylacetyl]pyrazin-2-yl}phenyl)methyl]-*N'*-(4-bromophenyl)urea: 1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 7.99 (s, 1H), 7.92-7.86 (m, 1H), 7.48-7.42 (m, 1H), 7.41-7.32 (m, 5H), 4.57-4.38 (m, 2H), 4.34-4.20 (m, 1H), 3.50-3.42 (m, 1H), 3.37-3.30 (m, 1H), 3.11-3.00 (m, 1H), 2.95-2.82 (m, 1H), 2.07-1.91 (m, 2H), 1.89-1.66 (m, 2H). MS (EI) for $C_{24}H_{26}N_7O_2Br$: 523 (MH^+).
- [1252] *N*-[(3-{5-Amino-6-[(3*R*)-piperidin-3-ylacetyl]pyrazin-2-yl}phenyl)methyl]-*N'*-naphthalen-2-ylurea: 1H NMR (400 MHz, d_4 -MeOH): δ 8.62 (s, 1H), 7.95-7.87 (m, 2H), 7.81-7.75 (m, 1H), 7.70-7.59 (m, 3H), 7.39-7.20 (m, 5H), 4.55-4.33 (m, 2H), 4.16-4.04 (m, 1H), 3.35-3.27 (m, 1H), 3.10-3.01 (m, 1H), 2.92-2.83 (m, 1H), 2.61-2.51 (m, 1H), 1.86-1.76 (m, 1H), 1.65-1.46 (m, 3H). MS (EI) for $C_{28}H_{29}N_7O_2$: 496 (MH^+).
- [1253] 3-Amino-6-[3-([(biphenyl-4-ylamino)carbonyl]amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.74 (s, 1H), 8.03 (s, 1H), 7.92-7.86 (m, 1H), 7.61-7.34 (m, 10H), 7.32-7.25 (m, 1H), 4.61-4.42 (m, 2H), 4.32-4.18 (m, 1H), 3.49-3.41 (m, 1H), 3.30-3.21 (m, 1H), 3.08-2.99 (m, 1H), 2.90-2.78 (m, 1H), 2.07-1.67 (m, 4H). MS (EI) for $C_{30}H_{31}N_7O_2$: 522 (MH^+).
- [1254] 3-Amino-6-[3-([(2,4-difluorophenyl)amino]carbonyl]amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.63 (s, 1H), 7.89 (s, 1H), 7.78-7.83 (m, 2H), 7.36 (t, 1H), 7.26-7.28 (m, 1H), 6.88-6.93 (m, 1H), 6.78-6.83 (m, 1H), 4.34-4.43 (m, 2H), 4.13-4.25 (m, 1H), 3.36-3.40 (m, 1H), 3.24-3.27

(m, 1H), 2.97-3.03 (m, 1H), 2.79-2.89 (m, 1H), 1.92-1.98 (m, 2H), 1.71-1.75 (m, 2H). MS (EI) for $C_{24}H_{25}N_7O_2F_2$: 482 (MH^+).

[1255] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-[3-({[(2-[(trifluoromethyl)oxy]phenyl} amino)carbonyl]amino)methyl}phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.76 (s, 1H), 8.13-8.15 (m, 1H), 7.99 (s, 1H), 7.90-7.92 (d, 1H), 7.46 (t, 1H), 7.36-7.38 (m, 1H), 7.26-7.30 (m, 2H), 7.04-7.09 (m, 1H), 4.44-4.55 (m, 2H), 4.02-4.03 (m, 1H), 3.45-3.48 (m, 1H), 3.07-3.13 (m, 2H), 2.92-2.95 (m, 1H), 1.77-2.05 (m, 4H). MS (EI) for $C_{25}H_{26}N_7O_3F_3$: 530 (MH^+).

[1256] 3-Amino-6-(3-({[(2-(methylthio)phenyl]amino)carbonyl]amino)methyl}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.64 (s, 1H), 7.93 (s, 1H), 7.80 (d, 1H), 7.69-7.71 (m, 1H), 7.26-7.38 (m, 3H), 7.09-7.14 (m, 1H), 6.96-6.98 (m, 1H), 4.34-4.46 (m, 2H), 4.09-4.13 (m, 1H), 3.31-3.38 (m, 1H), 2.90-2.98 (m, 2H), 2.78-2.83 (m, 1H), 2.80 (s, 3H), 1.84-1.94 (m, 2H), 1.66-1.73 (m, 2H). MS (EI) for $C_{25}H_{29}N_7O_2S$: 492 (MH^+).

[1257] 3-Amino-6-{3-({[(3-bromo-5-methylphenyl)amino]carbonyl]amino)methyl}phenyl}-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.73 (s, 1H), 8.01 (s, 1H), 7.89-7.91 (m, 1H), 7.54-7.57 (m, 1H), 7.45 (t, 1H), 7.34-7.37 (m, 2H), 7.26-7.29 (m, 1H), 4.42-4.54 (m, 2H), 4.21-4.30 (m, 1H), 3.45-3.48 (m, 1H), 2.86-3.15 (m, 3H), 2.21 (s, 3H), 1.99-2.02 (m, 2H), 1.78-1.82 (m, 2H). MS (EI) for $C_{25}H_{28}N_7O_2Br$: 539 (MH^+).

[1258] 3-Amino-6-{3-({[(2-bromophenyl)amino]carbonyl]amino)methyl}phenyl}-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.64 (s, 1H), 7.92 (s, 1H), 7.81-7.88 (m, 2H), 7.45-7.47 (m, 1H), 7.39 (t, 1H), 7.27-7.29 (m, 1H), 7.18-7.22 (m, 1H), 6.84-6.88 (m, 1H), 4.81 (s, 1H), 4.35-4.46 (m, 2H), 4.12-4.23 (m, 1H), 3.35-3.39 (m, 1H), 2.94-2.99 (m, 1H), 2.78-2.82 (m, 1H), 1.67-1.96 (m, 4H). MS (EI) for $C_{24}H_{26}N_7O_2Br$: 525 (MH^+).

[1259] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-[3-({[(4-[(trifluoromethyl)oxy]phenyl} amino)carbonyl]amino)methyl}phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.73 (s, 1H), 7.99 (s, 1H), 7.89-7.91 (m, 1H), 7.44-7.49 (m, 3H), 7.40-7.38 (m, 1H), 7.16-7.18 (m, 2H), 4.43-4.55 (m, 2H), 4.21-4.32 (m, 1H), 3.46-3.48 (m, 1H), 3.29-3.34 (m, 1H), 3.05-3.13 (m, 1H), 2.86-2.94 (m, 1H), 1.79-2.03 (m, 4H). MS (EI) for $C_{25}H_{26}N_7O_3F_3$: 530 (MH^+).

- [1260] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([4-(trifluoromethyl)phenyl]amino)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.73 (s, 1H), 8.00 (s, 1H), 7.89-7.91 (m, 1H), 7.53-7.60 (m, 4H), 7.44-7.47 (m, 1H), 7.36-7.38 (m, 1H), 4.44-4.56 (m, 2H), 4.22-4.33 (m, 1H), 3.43-3.50 (m, 1H), 3.04-3.12 (m, 2H), 2.83-2.94 (m, 1H), 1.73-2.08 (m, 4H). MS (EI) for $\text{C}_{25}\text{H}_{26}\text{N}_7\text{O}_2\text{F}_3$: 514 (MH^+).
- [1261] 3-Amino-6-[3-([(2,3-dihydro-1,4-benzodioxin-6-ylamino)carbonyl]amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.73 (s, 1H), 8.01 (s, 1H), 7.90-7.85 (m, 1H), 7.47-7.41 (m, 1H), 7.37-7.32 (m, 1H), 7.02-6.98 (m, 1H), 6.73-6.70 (m, 2H), 4.55-4.37 (m, 2H), 4.33-4.22 (m, 1H), 4.23-4.15 (m, 4H), 3.50-3.43 (m, 1H), 3.36-3.30 (m, 1H), 3.11-3.02 (m, 1H), 2.96-2.85 (m, 1H), 2.09-1.71 (m, 4H). MS (EI) for $\text{C}_{26}\text{H}_{29}\text{N}_7\text{O}_4$: 504 (MH^+).
- [1262] 3-Amino-6-[3-([(1,3-benzodioxol-5-ylamino)carbonyl]amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.71 (s, 1H), 8.01 (s, 1H), 7.90-7.85 (m, 1H), 7.47-7.41 (m, 1H), 7.37-7.32 (m, 1H), 7.10-7.07 (m, 1H), 6.74-6.64 (m, 2H), 5.90 (s, 2H), 4.54-4.38 (m, 2H), 4.33-4.22 (m, 1H), 3.50-3.43 (m, 1H), 3.37-3.31 (m, 1H), 3.12-3.03 (m, 1H), 2.96-2.86 (m, 1H), 2.08-1.96 (m, 2H), 1.89-1.72 (m, 2H). MS (EI) for $\text{C}_{25}\text{H}_{27}\text{N}_7\text{O}_4$: 490 (MH^+).
- [1263] 3-Amino-6-(3-([4-(methyloxy)phenyl]amino)carbonyl)amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 8.00 (s, 1H), 7.91-7.86 (m, 1H), 7.47-7.41 (m, 1H), 7.38-7.33 (m, 1H), 7.29-7.24 (m, 2H), 6.88-6.82 (m, 2H), 4.55-4.39 (m, 2H), 4.33-4.19 (m, 1H), 3.75 (s, 3H), 3.50-3.41 (m, 1H), 3.34-3.26 (m, 1H), 3.10-3.00 (m, 1H), 2.93-2.82 (m, 1H), 2.08-1.91 (m, 2H), 1.89-1.78 (m, 2H). MS (EI) for $\text{C}_{25}\text{H}_{29}\text{N}_7\text{O}_3$: 476 (MH^+).
- [1264] 3-Amino-6-[3-([4-(4-fluorophenyl)amino]carbonyl)amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.72 (s, 1H), 7.99 (s, 1H), 7.92-7.87 (m, 1H), 7.49-7.41 (m, 1H), 7.40-7.33 (m, 3H), 7.04-6.96 (m, 2H), 4.55-4.44 (m, 2H), 4.32-4.22 (m, 1H), 3.50-3.43 (m, 1H), 3.36-3.29 (m, 1H), 3.12-3.03 (m, 1H), 2.96-2.86 (m, 1H), 2.08-1.95 (m, 2H), 1.90-1.73 (m, 2H). MS (EI) for $\text{C}_{24}\text{H}_{26}\text{N}_7\text{O}_2\text{F}$: 464 (MH^+).
- [1265] 3-Amino-6-(3-([4-(dimethylamino)phenyl]amino)carbonyl)amino)methyl]phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -

MeOH): δ 8.72 (s, 1H), 7.99-7.90 (m, 2H), 7.65-7.58 (m, 2H), 7.54-7.42 (m, 3H), 7.39-7.34 (m, 1H), 4.49 (s, 2H), 4.36-4.24 (m, 1H), 3.53-3.44 (m, 1H), 3.39-3.30 (m, 1H), 3.15-3.05 (m, 1H), 3.04-2.90 (m, 1H), 2.15-1.97 (m, 2H), 1.95-1.82 (m, 2H). MS (EI) for $C_{26}H_{32}N_8O_2$: 489 (MH^+).

[1266] 3-Amino-6-(3-([(2,3-dihydro-1-benzofuran-5-ylsulfonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.65 (s, 1H), 7.93 (s, 1H), 7.89-7.83 (m, 1H), 7.63-7.55 (m, 2H), 7.39-7.33 (m, 1H), 7.24-7.19 (m, 1H), 6.77-6.72 (m, 1H), 4.62-4.53 (m, 2H), 4.39-4.24 (m, 1H), 4.15 (s, 2H), 3.54-3.46 (m, 1H), 3.39-3.31 (m, 1H), 3.21-3.10 (m, 3H), 3.06-2.92 (m, 1H), 2.19-2.03 (m, 2H), 1.96-1.81 (m, 2H). MS (EI) for $C_{25}H_{28}N_6O_4S$: 509 (MH^+).

[1267] 3-Amino-6-(3-([(3,4-dihydro-2*H*-1,5-benzodioxepin-7-ylsulfonyl)amino]methyl)phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.67 (s, 1H), 7.95 (s, 1H), 7.88-7.83 (m, 1H), 7.39-7.32 (m, 3H), 7.24-7.19 (m, 1H), 6.98-6.93 (m, 1H), 4.38-4.27 (m, 1H), 4.21-4.11 (m, 6H), 3.55-3.46 (m, 1H), 3.40-3.31 (m, 1H), 3.19-3.08 (m, 1H), 3.06-2.94 (m, 1H), 2.22-2.02 (m, 4H), 1.98-1.81 (m, 2H). MS (EI) for $C_{26}H_{30}N_6O_5S$: 539 (MH^+).

[1268] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([(quinolin-7-ylsulfonyl)amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.98-8.95 (m, 1H), 8.46 (s, 1H), 8.28-8.23 (m, 2H), 8.04-7.99 (m, 1H), 7.75 (s, 1H), 7.63-7.46 (m, 3H), 7.10-6.98 (m, 2H), 4.39-4.26 (m, 1H), 4.26 (s, 2H), 3.55-3.48 (m, 1H), 3.39-3.31 (m, 1H), 3.19-3.10 (m, 1H), 3.04-2.92 (m, 1H), 2.21-2.02 (m, 2H), 2.00-1.82 (m, 2H). MS (EI) for $C_{26}H_{27}N_7O_3S$: 518 (MH^+).

[1269] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([(2-(trifluoromethyl)phenyl)sulfonyl]amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.57 (s, 1H), 8.43 (d, 1H), 8.02-8.04 (m, 1H), 7.98 (s, 1H), 7.74-7.79 (m, 2H), 7.60-7.62 (m, 2H), 7.24-7.28 (m, 2H), 7.14 (d, 1H), 4.22 (s, 3H), 3.39-3.44 (m, 1H), 3.01-3.06 (m, 2H), 2.82-2.92 (m, 1H), 1.99-2.18 (m, 2H), 1.80-1.82 (m, 2H). MS (EI) for $C_{24}H_{25}N_6O_3F_3S$: 535 (MH^+).

[1270] 3-Amino-*N*-[(3*S*)-piperidin-3-yl]-6-(3-([(4-[(trifluoromethyl)oxy]phenyl)sulfonyl]amino]methyl)phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.58 (s, 1H), 8.46 (d, 1H), 7.89 (s, 1H), 7.76-7.84 (m, 2H), 7.24-7.30 (m, 3H), 7.12 (d, 1H), 4.15-4.27 (m, 1H), 4.13 (s, 2H), 3.39-3.43 (m, 1H), 3.01-3.07 (m, 2H), 2.91-

2.93 (m, 1H), 1.94-2.09 (m, 2H), 1.74-1.85 (m, 2H). MS (EI) for $C_{24}H_{25}N_6O_4F_3S$: 551 (MH^+).

[1271] 3-Amino-*N*-[(3S)-piperidin-3-yl]-6-(3-{[(2-[(trifluoromethyl)oxy]phenyl)sulfonyl]amino)methyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.67 (s, 1H), 8.02 (s, 1H), 7.94-8.02 (m, 1H), 7.83-7.85 (m, 1H), 7.60-7.64 (m, 1H), 7.38-7.41 (m, 2H), 7.31-7.35 (m, 1H), 7.21-7.23 (m, 1H), 4.29 (s, 3H), 3.47-3.53 (m, 1H), 3.11-3.16 (m, 2H), 2.96-3.10 (m, 1H), 1.84-2.38 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_4F_3S$: 551 (MH^+).

[1272] 3-Amino-*N*-[(3S)-piperidin-3-yl]-6-{3-[[4-(trifluoromethyl)phenyl]sulfonyl]amino)methyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.56 (s, 1H), 8.88 (s, 2H), 7.84 (m, 1H), 7.76 (d, 1H), 8.67 (d, 2H), 7.25 (t, 1H), 7.12 (d, 1H), 4.18-4.27 (m, 1H), 4.14 (s, 2H), 3.38-3.41 (m, 1H), 3.01-3.06 (m, 2H), 2.87-2.92 (m, 1H), 1.80-2.06 (m, 4H). MS (EI) for $C_{24}H_{25}N_6O_3F_3S$: 535 (MH^+).

[1273] 3-Amino-6-(3-{[(2,3-dihydro-1,4-benzodioxin-6-ylsulfonyl)amino)methyl}phenyl)-*N*-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.67 (s, 1H), 7.96 (s, 1H), 7.88-7.84 (m, 1H), 7.39-7.33 (m, 1H), 7.30-7.25 (m, 2H), 7.24-7.19 (m, 1H), 6.90-6.86 (m, 1H), 4.38-4.27 (m, 1H), 4.26-4.19 (m, 4H), 4.16 (s, 2H), 3.54-3.46 (m, 1H), 3.39-3.31 (m, 1H), 3.18-3.10 (m, 1H), 3.05-2.95 (m, 1H), 2.20-2.05 (m, 2H), 1.97-1.83 (m, 2H). MS (EI) for $C_{25}H_{28}N_6O_5S$: 524 (MH^+).

[1274] 3-Amino-*N*-[(3S)-piperidin-3-yl]-6-(3-{[(2-thienylsulfonyl)amino)methyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.07 (s, 1H), 7.91-7.86 (m, 1H), 7.77-7.74 (m, 1H), 7.63-7.59 (m, 1H), 7.42-7.36 (m, 1H), 7.28-7.23 (m, 1H), 7.14-7.10 (m, 1H), 4.37-4.25 (m, 1H), 4.23 (s, 2H), 3.54-3.45 (m, 1H), 3.39-3.31 (m, 1H), 3.19-3.08 (m, 1H), 3.05-2.94 (m, 1H), 2.19-2.02 (m, 2H), 1.97-1.81 (m, 2H). MS (EI) for $C_{21}H_{24}N_6O_3S_2$: 473 (MH^+).

[1275] 3-Amino-*N*-[(3S)-piperidin-3-yl]-6-(3-{[(3-thienylsulfonyl)amino)methyl}phenyl)pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 8.09-8.03 (m, 2H), 7.91-7.86 (m, 1H), 7.60-7.55 (m, 1H), 7.41-7.33 (m, 2H), 7.27-7.21 (m, 1H), 4.37-4.25 (m, 1H), 4.20 (s, 2H), 3.54-3.45 (m, 1H), 3.39-3.31 (m, 1H), 3.19-3.09 (m, 1H), 3.05-2.93 (m, 1H), 2.19-2.02 (m, 2H), 1.98-1.81 (m, 2H). MS (EI) for $C_{21}H_{24}N_6O_3S_2$: 473 (MH^+).

- [1276] 3-Amino-6-[3-([(4-chlorophenyl)sulfonyl]amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_4 -MeOH): δ 8.66 (s, 1H), 7.94 (s, 1H), 7.90-7.85 (m, 1H), 7.81-7.75 (m, 2H), 7.50-7.45 (m, 2H), 7.39-7.33 (m, 1H), 7.25-7.20 (m, 1H), 4.38-4.25 (m, 1H), 4.20 (s, 2H), 3.55-3.46 (m, 1H), 3.39-3.31 (m, 1H), 3.18-3.08 (m, 1H), 3.05-2.93 (m, 1H), 2.20-2.01 (m, 2H), 1.98-1.81 (m, 2H). MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_6\text{O}_3\text{SCl}$: 501 (MH^+).
- [1277] N-[(2,4-Dichlorophenyl)methyl]-N'-[(1S)-2,3-dihydro-1H-inden-1-yl]benzene-1,3-dicarboxamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.20 (t, $J = 6.0$ Hz, 1H), 8.93 (d, $J = 8.4$ Hz, 1H), 8.43 (m, 1H), 8.05 (m, 2H), 7.64 (m, 1H), 7.59 (m, 1H), 7.42 (m, 2H), 7.25 (m, 4H), 5.59 (m, 1H), 4.53 (m, 2H), 3.00 (m, 1H), 2.86 (m, 1H), 2.47 (m, 1H), 2.01 (m, 1H). MS (EI) for $\text{C}_{24}\text{H}_{20}\text{Cl}_2\text{N}_2\text{O}_2$: 439 (MH^+).

Example 64

3-Amino-6-{3-[(3-fluoro-5-(trifluoromethyl)phenyl)methyl]amino}carbonyl phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

- [1278] To a solution of 3-[5-amino-6-((3S)-1-(tert-butoxycarbonyl)piperidine-3-yl)amino]carbonylpyrazine-2-yl]benzoic acid (80 mg, 0.18 mmol) in dry dichloromethane (1.0 mL) were added 1-ethyl-(3-dimethylaminopropyl)carbodiimide hydrochloride (EDC, 48 mg, 0.25 mmol) and 1-hydroxybenzotriazole (HOBt, 34 mg, 0.25 mmol). The reaction mixture was stirred for 10-15 min at room temperature. A solution of 3-fluoro-4-(trifluoromethyl)benzylamine (52 mg, 0.27 mmol) in dichloromethane (1 mL) was added and stirring continued for 2-3 h. Reaction was diluted with ethyl acetate (150 mL), washed with saturated aqueous sodium bicarbonate (40 mL) and saturated aqueous sodium chloride (50 mL). Dried over anhydrous magnesium sulfate, filtered, concentrated, and purification on silica (7:3 hexanes/ethyl acetate) afforded the amide product. MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2\text{F}_4$: 517 [$(\text{M-boc})\text{H}^+$].
- [1279] To a solution of the above amide in methanol (2 mL) was added 4.0 N hydrochloric acid (1,4-dioxane solution, 1 mL) and the reaction stirred until the deprotection (removal of Boc-protecting group) was completed as monitored by thin layer chromatography. The excess reagent and solvent was removed at reduced pressure to give a solid product which was washed further with ethyl acetate (50 mL) to afford the hydrochloric acid salt of 3-amino-6-{3-[(3-fluoro-5-(trifluoromethyl)phenyl)methyl]}

amino)carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide (16 mg, 15% yield) as yellow solid. ¹H NMR (400 MHz, d₆-DMSO): 9.90 (s, 1H), 9.25 (br s, 1H), 9.02 (m, 3H), 8.80 (s, 1H), 8.54 (d, 1H), 7.90 (d, 1H), 7.60 (m, 4H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.70 (m, 1H), 1.92 (m, 4H); MS (EI) for C₂₅H₂₄N₆O₂F₄: 517 (MH⁺).

[1280] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[1281] 3-amino-6-[3-({[(3,4-dichlorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.80 (t, 1H), 9.20 (br s, 1H), 8.93 (m, 3H), 8.70 (s, 1H), 8.58 (d, 1H), 7.90 (d, 1H), 7.60 (m, 3H), 7.40 (d, 1H), 4.50 (d, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 1.90 (m, 4H); MS (EI) for C₂₄H₂₄N₆O₂Cl₂: 499, 501 (MH⁺).

[1282] 3-amino-6-[3-({[(5-chloro-2-methylphenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.45 (t, 1H), 9.20 (br s, 1H), 9.01 (m, 3H), 8.70 (s, 1H), 8.35 (d, 1H), 7.90 (d, 1H), 7.30 (s, 1H), 7.20 (s, 2H), 4.50 (m, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.70 (m, 1H), 2.38 (s, 3H), 1.90 (m, 4H); MS (EI) for C₂₅H₂₇N₆O₂Cl: 479, 480 (MH⁺).

[1283] 3-amino-6-[3-({[(2-chloro-5-(trifluoromethyl)phenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.85 (t, 1H), 9.25 (br s, 1H), 9.02 (m, 3H), 8.80 (s, 1H), 8.40 (d, 1H), 7.90 (d, 1H), 7.70 (m, 3H), 7.60 (t, 1H), 4.64 (br s, 2H), 4.30 (m, 1H), 3.25 (m, 3H), 2.70 (m, 1H), 1.92 (m, 4H); MS (EI) for C₂₅H₂₄N₆O₂F₃Cl: 533, 534 (MH⁺).

[1284] 3-amino-6-[3-({[(2-chloro-6-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.30 (br s, 1H), 9.00 (br s, 1H), 9.01 (t, 1H), 8.95 (s, 1H), 8.80 (d, 1H), 8.73 (s, 1H), 8.30 (d, 1H), 7.85 (d, 1H), 7.60 (t, 1H), 7.40 (m, 2H), 7.30 (t, 1H), 4.60 (s, 2H), 4.45 (m, 1H), 3.30 (m, 1H), 3.12 (m, 1H), 2.70 (m, 1H), 1.92 (m, 4H); MS (EI) for C₂₄H₂₄N₆O₂FCl: 483 (MH⁺).

[1285] 3-amino-6-[3-({[(3-chloro-2-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): 9.80 (t, 1H), 9.30 (br s, 1H), 9.02 (br s, 1H), 8.92 (m, 2H), 8.70 (s, 1H), 8.30 (d, 1H), 7.90 (d, 1H), 7.60 (t, 1H), 7.50 (m, 2H), 7.20 (t, 1H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.90 (m, 1H), 1.92 (m, 4H); MS (EI) for C₂₄H₂₄N₆O₂FCl: 483 (MH⁺).

[1286] 3-amino-6-[3-([2-fluoro-5-(trifluoromethyl)phenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.65 (t, 1H), 9.20 (br s, 1H), 9.00 (br s, 1H), 8.90 (m, 2H), 8.70 (s, 1H), 8.30 (d, 1H), 7.90 (d, 1H), 7.80 (m, 1H), 7.78 (m, 1H), 7.58 (t, 1H), 7.45 (t, 1H), 4.60 (br s, 2H), 4.32 (m, 1H), 3.20 (m, 3H), 2.70 (m, 1H), 1.92 (m, 4H); MS (EI) for $\text{C}_{25}\text{H}_{24}\text{N}_6\text{O}_2\text{F}_4$: 517 (MH^+).

Example 65

3-amino-6-[3-([3-chloro-2,6-difluorophenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

[1287] To a solution of tert-butyl (3S)-3-([3-amino-6-[3-(methoxycarbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]piperidine-1-carboxylate (100 mg, 0.22 mmol) in tetrahydrofuran/water (4:1, 1.5 mL) was added 0.5 N aqueous lithium hydroxide (0.5 mL, 0.55 mmol) and the reaction was stirred at room temperature overnight. Reaction was diluted with dichloromethane (4 mL) and treated with 0.5 N aqueous hydrochloric acid (0.5 mL). The aqueous layer was discarded. The crude acid in dichloromethane solution was treated with 1-ethyl-(3-dimethylaminopropyl)carbodiimide hydrochloride (EDCI, 63 mg, 0.33 mmol) and 1-hydroxybenzotriazole (HOBT, 45 mg, 0.33 mmol). The reaction was stirred for 10-15 min and then a solution of 5-chloro-1,6-difluorobenzylamine (46 mg, 0.26 mmol) in dichloromethane (0.5 mL) was added. The reaction was stirred for 2-3 h and diluted with ethyl acetate (150 mL). The organic layer was washed with saturated aqueous sodium bicarbonate (40 mL), saturated aqueous sodium chloride (50 mL), and dried over anhydrous magnesium sulfate. Filtration, concentration, and purification on silica (3:7 hexanes/ethyl acetate) afforded the boc-protected intermediate. MS (EI) for $\text{C}_{24}\text{H}_{23}\text{N}_6\text{O}_2\text{F}_2\text{Cl}$: 501, 503 [$(\text{M-boc})\text{H}^+$].

[1288] To a solution of the above intermediate in methanol (2 mL) was added 4.0 N hydrochloric acid (1,4-dioxane solution, 1 mL) and the reaction stirred until the deprotection was completed as monitored by thin layer chromatography. The excess reagent and solvent was removed at reduced pressure to give a solid product which was washed further with ethyl acetate (50 mL) to afford the hydrochloric acid salt of 3-amino-6-[3-([3-chloro-2,6-difluorophenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide (76 mg, 69% yield) as yellow solid ^1H NMR (400 MHz,

d_6 -DMSO): 9.26 (m, 2H), 9.10 (m, 1H), 8.90 (s, 1H), 8.80 (d, 1H), 8.60 (s, 1H), 8.35 (d, 1H), 7.60 (m, 2H), 7.20 (t, 1H), 7.20 (t, 1H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 3.10 (m, 1H), 2.80 (m, 1H), 1.90 (m, 2H), 1.80 (m, 2H); MS (EI) for $C_{24}H_{23}N_6O_2F_2Cl$: 501, 503 (MH^+).

[1289] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[1290] 3-amino-6-[3-([2-fluoro-4-(trifluoromethyl)phenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.70 (t, 1H), 9.30 (br s, 1H), 9.10 (br s, 1H), 9.01 (m, 2H), 8.80 (s, 1H), 8.36 (d, 1H), 7.90 (d, 1H), 7.60 (m, 6H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.78 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{25}H_{24}N_6O_2F_4$: 517 (MH^+).

[1291] 3-amino-6-[3-([2-chloro-4-fluorophenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.54 (m, 1H), 9.25 (br s, 1H), 9.02 (m, 3H), 8.74 (s, 1H), 8.34 (d, 1H), 7.90 (d, 1H), 7.50 (m, 4H), 7.20 (t, 1H), 4.58 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.75 (m, 1H), 1.92 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2F_2Cl$: 483, 485 (MH^+).

[1292] 3-amino-6-[3-([2,3-dichlorophenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.80 (t, 1H), 9.20 (br s, 1H), 9.01 (m, 3H), 8.74 (s, 1H), 8.30 (d, 1H), 7.90 (d, 1H), 7.60 (m, 2H), 7.40 (m, 2H), 4.60 (d, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.74 (m, 1H), 1.92 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2Cl_2$: 499, 500 (MH^+).

[1293] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-([2,3,6-trifluorophenyl]methyl)amino]carbonylphenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.23 (m, 2H), 9.02 (br s, 1H), 8.90 (s, 1H), 8.80 (d, 1H), 8.32 (d, 1H), 7.60 (t, 1H), 7.50 (m, 1H), 7.20 (m, 1H), 4.60 (d, 2H), 4.34 (m, 1H), 3.30 (m, 2H), 3.10 (m, 1H), 2.80 (m, 1H), 1.90 (m, 2H), 1.80 (m, 2H); MS (EI) for $C_{24}H_{23}N_6O_2F_3$: 485 (MH^+).

[1294] 3-amino-6-[3-([4-chloro-3-(trifluoromethyl)phenyl]methyl)amino]carbonylphenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.80 (m, 2H), 9.30 (br s, 1H), 9.04 (m, 3H), 8.74 (s, 1H), 8.30 (d, 1H), 7.90 (m, 2H), 7.70 (s, 2H), 7.60 (t, 1H), 4.60 (d, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{25}H_{24}N_6O_2F_3Cl$: 533, 534 (MH^+).

- [1295] 3-amino-6-[3-({[(3-chloro-4-fluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.82 (m, 1H), 9.30 (br s, 1H), 9.02 (m, 3H), 8.78 (s, 1H), 8.50 (d, 1H), 7.90 (m, 1H), 7.70 (s, 2H), 7.60 (m, 2H), 7.40 (m, 2H), 4.50 (d, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 1.92 (m, 4H); MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2\text{FCl}$: 483, 485 (MH^+).
- [1296] 3-amino-6-[3-({[(2,6-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.20 (br s, 1H), 9.10 (m, 1H), 8.95 (m, 2H), 8.80 (d, 1H), 8.74 (s, 1H), 8.30 (d, 1H), 7.84 (d, 1H), 7.60 (t, 1H), 7.40 (m, 1H), 7.10 (m, 2H), 4.60 (d, 2H), 4.30 (m, 1H), 3.30 (m, 2H), 3.06 (m, 1H), 2.78 (m, 1H), 1.90 (m, 2H), 1.74 (m, 2H); MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2\text{F}$: 467 (MH^+).
- [1297] 3-amino-6-[3-({[(3,4-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.80 (m, 1H), 9.20 (br s, 1H), 8.99 (m, 3H), 8.70 (s, 1H), 8.30 (d, 1H), 7.90 (d, 1H), 7.60 (t, 1H), 7.40 (m, 2H), 7.20 (br s, 1H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 1.92 (m, 4H); MS (EI) for $\text{C}_{24}\text{H}_{24}\text{N}_6\text{O}_2\text{F}_2$: 467 (MH^+).
- [1298] 3-amino-6-[3-({[(6-chloro-2-fluoro-3-methylphenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.20 (br s, 1H), 8.90 (m, 3H), 8.75 (d, 1H), 8.50 (s, 1H), 8.30 (d, 1H), 7.86 (d, 1H), 7.60 (t, 1H), 7.40 (m, 2H), 7.20 (t, 1H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.30 (m, 2H), 3.02 (m, 1H), 2.80 (m, 1H), 2.30 (s, 3H), 1.92 (m, 2H), 1.80 (m, 2H); MS (EI) for $\text{C}_{25}\text{H}_{26}\text{N}_6\text{O}_2\text{F}_1\text{Cl}$: 497, 499 (MH^+).
- [1299] 3-amino-6-[3-({[(2-chloro-6-fluoro-3-methylphenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.30 (m, 1H), 9.10 (m, 1H), 8.90 (m, 2H), 8.80 (d, 1H), 8.70 (s, 1H), 8.30 (d, 1H), 7.84 (d, 1H), 7.60 (t, 1H), 7.30 (m, 2H), 4.60 (br s, 2H), 4.30 (m, 1H), 3.30 (m, 2H), 3.02 (m, 1H), 2.80 (m, 1H), 2.30 (s, 3H), 1.92 (m, 2H), 1.74 (m, 2H); MS (EI) for $\text{C}_{25}\text{H}_{26}\text{N}_6\text{O}_2\text{F}_1\text{Cl}$: 497, 499 (MH^+).
- [1300] 3-amino-6-[3-({[(5-fluoro-2-methylphenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, d_6 -DMSO): 9.40 (m, 1H), 9.20 (br s, 1H), 8.94 (s, 1H), 8.85 (d, 1H), 8.72 (s, 1H), 8.57 (d, 1H), 7.90 (d, 1H), 7.60 (t, 1H), 7.24 (m, 1H), 7.15 (m, 1H), 7.02 (m, 1H), 4.50 (br s, 2H), 4.30 (m, 1H),

3.20 (m, 3H), 2.80 (m, 1H), 2.30 (s, 3H), 1.90 (m, 4H); MS (EI) for $C_{25}H_{27}N_6O_2F$: 463 (MH^+).

[1301] 3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.50 (m, 1H), 9.10 (br s, 1H), 8.94 (s, 1H), 8.90 (d, 1H), 8.70 (s, 1H), 8.40 (d, 1H), 7.90 (d, 1H), 7.60 (t, 1H), 7.40 (s, 1H), 7.25 (m, 2H), 4.50 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 2.30 (s, 3H), 1.90 (m, 4H); MS (EI) for $C_{25}H_{27}N_6O_2FCl$: 479, 481 (MH^+).

[1302] 3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.30 (br s, 1H), 9.10 (br s, 1H), 8.90 (s, 1H), 8.84 (d, 1H), 8.63 (s, 1H), 8.30 (m, 2H), 7.84 (d, 1H), 7.60 (t, 1H), 7.30 (t, 1H), 6.70 (d, 2H), 4.45 (br s, 2H), 4.20 (m, 1H), 3.80 (s, 6H), 3.20 (m, 2H), 2.90 (m, 1H), 2.70 (s, 1H), 1.80 (m, 4H); MS (EI) for $C_{26}H_{30}N_6O_4$: 491 (MH^+).

[1303] 3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.80 (m, 1H), 9.30 (br s, 1H), 9.01 (m, 3H), 8.85 (s, 1H), 8.34 (d, 1H), 7.90 (d, 1H), 7.70 (d, 1H), 7.60 (t, 1H), 7.40 (m, 2H), 4.50 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2FBr$: 427, 429 (MH^+).

[1304] 3-amino-6-[3-(((3-bromo-5-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.50 (m, 1H), 9.20 (br s, 1H), 8.98 (m, 2H), 8.70 (s, 1H), 8.36 (d, 1H), 7.90 (d, 1H), 7.60 (m, 3H), 7.20 (t, 1H), 4.50 (br s, 2H), 4.30 (m, 1H), 3.20 (m, 3H), 2.70 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2FBr$: 527, 529 (MH^+).

[1305] 3-amino-6-[3-(((3-methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_6 -DMSO): 9.50 (t, 1H), 9.30 (br s, 1H), 8.90 (m, 2H), 8.70 (s, 1H), 8.35 (d, 1H), 7.90 (d, 1H), 7.60 (m, 3H), 7.22 (t, 1H), 6.93 (br s, 1H), 6.80 (d, 1H), 4.50 (br s, 2H), 4.30 (m, 1H), 3.73 (s, 3H), 3.20 (m, 3H), 2.70 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{25}H_{28}N_6O_3$: 461 (MH^+).

[1306] 3-amino-6-[3-(((3,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, $DMSO-d_6$): 9.30 (s, 1H), 9.20 (br s, 1H), 8.95 (m, 2H), 8.86 (d, 1H), 8.60 (s, 1H), 8.30 (d, 1H), 7.86 (d, 1H), 7.58 (t, 1H), 7.02 (s, 1H), 6.87 (br s, 2H), 4.50 (d, 2H), 4.30 (m, 1H), 3.74 (s, 3H), 3.72 (s,

3H), 3.25 (m, 2H), 3.15 (m, 1H), 2.70 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{26}H_{30}N_6O_4$: 491 (MH^+).

[1307] 3-amino-6-[3-({[(2-chloro-3,6-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): 9.31 (s, 1H), 9.09 (s, 2H), 8.93 (s, 1H), 8.80 (d, 1H), 8.56 (s, 1H), 8.33 (d, 1H), 7.86 (d, 1H), 7.57 (t, 1H), 7.48 (m, 1H), 7.35 (m, 1H), 4.65 (m, 2H), 4.37 (m, 1H), 3.29 (m, 2H), 3.05 (m, 1H), 2.74 (m, 1H), 1.99-1.72 (m, 4H); MS (EI) for $C_{24}H_{23}N_6O_2F_2Cl$: 501.04 (M^+).

[1308] 3-amino-6-[3-({[(2,5-dichlorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): 9.61 (t, 1H), 9.15 (m, 1H), 8.97 (s, 1H), 8.96 (s, 1H), 8.94 (s, 1H), 8.77 (s, 1H), 8.36 (d, 1H), 7.94 (d, 1H), 7.61 (t, 1H), 7.53 (d, 1H), 7.44-7.39 (m, 2H), 4.57 (m, 2H), 4.29 (m, 1H), 3.28-3.16 (m, 3H), 2.76 (m, 1H), 1.91-1.72 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2Cl_2$: 500.98 (MH^+).

[1309] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-({[(2,3,4-trifluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): 9.56 (s, 1H), 9.18 (s, 1H), 9.00 (s, 1H), 8.96 (s, 1H), 8.92 (d, 1H), 8.70 (s, 1H), 8.34 (d, 1H), 7.89 (d, 1H), 7.59 (t, 1H), 7.32 (m, 2H), 4.55 (d, 2H), 4.28 (m, 1H), 3.29-3.15 (m, 3H), 2.76 (m, 1H), 1.91-1.73 (m, 4H); MS (EI) for $C_{24}H_{23}N_6O_2F_3$: 485.06 (MH^+).

[1310] 3-amino-6-[3-({[(2,3-difluorophenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): 9.60 (s, 1H), 9.28 (s, 1H), 9.06 (s, 1H), 8.96 (s, 1H), 8.94 (s, 1H), 8.73 (s, 1H), 8.34 (d, 1H), 7.90 (d, 1H), 7.59 (t, 1H), 7.37-7.16 (m, 3H), 4.60 (m, 2H), 4.30 (m, 1H), 3.27-3.15 (m, 3H), 2.77 (m, 1H), 1.92-1.73 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2F_2$: 467.06 (MH^+).

[1311] 3-amino-6-[3-({[(3-fluoro-4-methylphenyl)methyl]amino}carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): 9.55 (s, 1H), 9.28 (s, 1H), 9.08 (s, 1H), 8.96 (m, 2H), 8.72 (s, 1H), 8.33 (d, 1H), 7.91 (d, 1H), 7.59 (t, 1H), 7.24 (t, 1H), 7.15-7.10 (m, 2H), 4.50 (m, 2H), 4.30 (m, 1H), 3.29-3.13 (m, 3H), 2.76 (m, 1H), 2.20 (s, 3H), 1.94-1.71 (m, 4H); MS (EI) for $C_{25}H_{27}N_6O_2F$: 463.12 (MH^+).

[1312] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-({[(3,4,5-trifluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): 9.72 (m, 1H), 9.23 (s, 1H), 9.00 (m, 2H), 8.96 (s, 1H), 8.78 (s, 1H), 8.33 (d, 1H), 7.91 (d, 1H), 7.59 (t, 1H), 7.36 (m, 2H), 4.49 (m, 2H), 4.30 (m, 1H), 3.28-3.17 (m, 3H), 2.76 (m, 1H), 1.94-1.70 (m, 4H); MS (EI) for $C_{24}H_{23}N_6O_2F_3$: 485.28 (MH^+).

- [1313] 3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.51 (m, 1H), 9.29 (s, 1H), 9.08 (s, 1H), 8.94 (m, 2H), 8.70 (s, 1H), 8.33 (d, 1H), 7.91 (d, 1H), 7.58 (t, 1H), 7.25 (m, 2H), 7.09 (t, 1H), 4.46 (m, 2H), 4.29 (m, 1H), 3.30-3.13 (m, 3H), 2.76 (m, 1H), 2.22 (s, 1H), 1.94-1.70 (m, 4H); MS (EI) for $\text{C}_{25}\text{H}_{27}\text{N}_6\text{O}_2\text{F}$: 463.31 (MH^+).
- [1314] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.67 (s, 2H), 9.21 (s, 1H), 8.99 (s, 2H), 8.75 (m, 1H), 8.34 (d, 1H), 7.90 (d, 1H), 7.59 (t, 1H), 7.46 (m, 1H), 7.16 (m, 1H), 4.58 (m, 2H), 4.29 (m, 1H), 3.22 (m, 3H), 2.75 (m, 1H), 1.95-1.68 (m, 4H); MS (EI) for $\text{C}_{24}\text{H}_{23}\text{N}_6\text{O}_2\text{F}_3$: 485.27 (MH^+).
- [1315] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.63 (s, 1H), 9.36 (s, 1H), 9.14 (s, 1H), 8.98 (m, 2H), 8.76 (s, 1H), 8.48 (d, 1H), 8.34 (d, 1H), 7.93 (d, 1H), 7.81 (d, 2H), 7.73 (d, 1H), 7.59 (t, 1H), 7.51 (d, 2H), 6.54 (m, 1H), 4.56 (m, 2H), 4.31 (m, 1H), 3.32-3.12 (m, 3H), 2.75 (m, 1H), 1.94-1.69 (m, 4H); MS (EI) for $\text{C}_{27}\text{H}_{28}\text{N}_8\text{O}_2$: 497.06 (MH^+).
- [1316] 3-amino-6-(3-(((biphenyl-3-yl)methyl)amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.60 (t, 1H), 9.31 (s, 1H), 9.08 (s, 1H), 8.94 (m, 2H), 8.74 (s, 1H), 8.34 (d, 1H), 7.93 (d, 1H), 7.68-7.34 (m, 10H), 4.60 (m, 2H), 4.29 (m, 1H), 3.30-3.11 (m, 3H), 2.74 (m, 1H), 1.94-1.67 (m, 4H); MS (EI) for $\text{C}_{30}\text{H}_{30}\text{N}_6\text{O}_2$: 507.07 (MH^+).
- [1317] 3-amino-6-[3-(((2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.33 (t, 2H), 9.11 (m, 1H), 8.96 (s, 1H), 8.92 (d, 1H), 8.71 (s, 1H), 8.34 (d, 1H), 7.92 (d, 1H), 7.59 (t, 1H), 7.32 (m, 1H), 7.17 (m, 3H), 4.50 (m, 2H), 4.29 (m, 1H), 3.30-3.06 (m, 3H), 2.74 (m, 1H), 2.36 (s, 1H), 1.94-1.66 (m, 4H); MS (EI) for $\text{C}_{25}\text{H}_{28}\text{N}_6\text{O}_2$: 445.08 (MH^+).
- [1318] 3-amino-6-[3-(((2,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.34 (s, 1H), 9.28 (m, 1H), 9.13 (s, 1H), 8.96 (s, 1H), 8.91 (d, 1H), 8.69 (s, 1H), 8.34 (d, 1H), 7.92 (d, 1H), 7.59 (t, 1H), 7.11 (s, 1H), 7.06 (d, 1H), 6.97 (d, 1H), 4.46 (m, 2H), 4.29 (m, 1H), 3.32-3.02 (m, 3H), 2.74 (m, 1H), 2.31 (s, 3H), 2.24 (s, 3H), 1.97-1.67 (m, 4H); MS (EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}_2$: 459.09 (MH^+).

- [1319] 3-amino-6-[3-([(3,5-dimethylphenyl)methyl]amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.39 (s, 1H), 9.28 (s, 1H), 9.06 (s, 1H), 8.95 (s, 1H), 8.91 (d, 1H), 8.68 (s, 1H), 8.34 (d, 1H), 7.92 (d, 1H), 7.59 (t, 1H), 6.96 (s, 2H), 6.87 (s, 1H), 4.44 (m, 2H), 4.28 (m, 1H), 3.31-3.08 (m, 3H), 2.74 (m, 1H), 2.31 (s, 6H), 1.95-1.66 (m, 4H); MS (EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}_2$: 459.09 (MH^+).
- [1320] 3-amino-6-[3-([(2,4-dimethylphenyl)methyl]amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.29 (s, 1H), 9.23 (t, 1H), 9.08 (m, 1H), 8.95 (s, 1H), 8.91 (d, 1H), 8.66 (s, 1H), 8.33 (d, 1H), 7.91 (d, 1H), 7.58 (t, 1H), 7.20 (d, 1H), 6.99 (s, 1H), 6.96 (d, 1H), 4.45 (m, 2H), 4.28 (m, 1H), 3.27-3.05 (m, 3H), 2.74 (m, 1H), 2.32 (s, 3H), 2.24 (s, 3H), 1.96-1.67 (m, 4H); MS (EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}_2$: 459.09 (MH^+).
- [1321] 3-amino-6-[3-([(phenyl)methyl]amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.48 (t, 1H), 9.26 (m, 1H), 9.05 (m, 1H), 8.96 (s, 1H), 8.92 (d, 1H), 8.70 (s, 1H), 8.34 (d, 1H), 7.91 (d, 1H), 7.59 (t, 1H), 7.40-7.31 (m, 4H), 7.28-7.22 (m, 1H), 4.53 (m, 2H), 4.29 (m, 1H), 3.32-3.07 (m, 3H), 2.75 (m, 1H), 1.95-1.67 (m, 4H); MS (EI) for $\text{C}_{24}\text{H}_{26}\text{N}_6\text{O}_2$: 431.09 (MH^+).
- [1322] 3-amino-6-[3-([(3,5-bis(methyloxy)phenyl)methyl]amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ^1H NMR (400 MHz, DMSO- d_6): 9.46 (t, 1H), 9.31 (m, 1H), 9.10 (m, 1H), 8.96 (s, 1H), 8.94 (d, 1H), 8.70 (s, 1H), 8.34 (d, 1H), 7.91 (d, 1H), 7.59 (t, 1H), 6.54 (d, 2H), 6.39 (t, 1H), 4.45 (m, 2H), 4.29 (m, 1H), 3.72 (s, 6H), 3.34-3.10 (m, 3H), 2.75 (m, 1H), 1.96-1.66 (m, 4H); MS (EI) for $\text{C}_{26}\text{H}_{30}\text{N}_6\text{O}_4$: 491.06 (MH^+).

Example 66

3-amino-6-(3-([(2-methylphenyl)acetyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

- [1323] To a solution of 2-methylphenylacetic acid (45 mg, 0.30 mmol) in dry dichloromethane (1.0 mL) were added 1-ethyl-(3-dimethylaminopropyl)carbodiimide hydrochloride (EDCI, 72 mg, 0.38 mmol) and 1-hydroxybenzotriazole (HOBT, 51 mg, 0.38 mmol). The reaction mixture was stirred for 10-15 min at room temperature. A solution of tert-butyl (3S)-3-([(3-amino-6-(3-aminophenyl)pyrazin-2-

yl]carbonyl}amino)piperidine-1-carboxylate (80 mg, 0.20 mmol) in dichloromethane (1 mL) was added and stirring continued for 2-3 h. Reaction was diluted with ethyl acetate (150 mL), washed with saturated aqueous sodium bicarbonate (40 mL) and saturated aqueous sodium chloride (50 mL). Dried over anhydrous magnesium sulfate, filtered, concentrated, and purification on silica (hexanes/ethyl acetate, 3:7) afforded the protected intermediate. MS (EI) for $C_{25}H_{28}N_6O_2$: 445 [(M-boc)H⁺].

[1324] To a solution of the above intermediate in methanol (2 mL) was added 4.0 N hydrochloric acid (1,4-dioxane solution, 1 mL) and the reaction stirred until the deprotection was completed as monitored by thin layer chromatography. The excess reagent and solvent was removed at reduced pressure to give a solid product which was washed further with ethyl acetate (50 mL) to afford the hydrochloric acid salt of 3-amino-6-(3-[(2-methylphenyl)acetyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide (56 mg, 63% yield) as yellow solid. ¹NMR (400 MHz, d₆-DMSO): 10.72 (s, 1H), 9.20 (br s, 1H), 9.04 (br s, 1H), 8.80 (s, 1H), 8.70 (d, 1H), 8.45 (s, 1H), 7.80 (d, 1H), 7.40 (m, 2H), 7.18 (m, 3H), 4.32 (m, 1H), 3.80 (s, 2H), 3.25 (m, 3H), 2.80 (m, 1H), 2.3 (s, 3H), 1.80 (m, 4H); MS (EI) for $C_{25}H_{28}N_6O_2$: 445 (MH⁺).

[1325] Using the same or analogous synthetic techniques and/or substituting with alternative reagents as described above, the following compounds of the invention were prepared:

[1326] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-({[3-(trifluoromethyl)phenyl]acetyl}amino)phenyl]pyrazine-2-carboxamide: ¹NMR (400 MHz, d₆-DMSO): 10.82 (s, 1H), 9.10 (br s, 1H), 8.92 (br s, 1H), 8.90 (s, 1H), 8.85 (d, 1H), 8.40 (s, 1H), 7.72 (m, 6H), 7.40 (t, 1H), 4.40 (m, 1H), 3.84 (s, 2H), 3.25 (m, 3H), 2.80 (m, 1H), 1.84 (m, 4H); MS (EI) for $C_{25}H_{25}N_6O_2F_3$: 499 (MH⁺).

[1327] 3-amino-6-[3-({[4-(methoxy)phenyl]acetyl}amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹NMR (400 MHz, d₆-DMSO): 10.60 (s, 1H), 9.10 (br s, 1H), 9.01 (br s, 1H), 8.78 (s, 1H), 8.68 (d, 1H), 8.40 (s, 1H), 7.80 (t, 1H), 7.40 (t, 1H), 7.30 (d, 2H), 6.85 (d, 2H), 4.45 (m, 1H), 3.72 (s, 3H), 3.60 (s, 2H), 3.25 (m, 3H), 2.80 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{25}H_{28}N_6O_3$: 461 (MH⁺).

[1328] 3-amino-6-(3-[(6-methylpyridin-3-yl)carbonyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹NMR (400 MHz, d₆-DMSO): 11.40 (s, 1H), 9.60 (s, 1H), 9.50 (br s, 1H), 9.10 (m, 2H), 8.90 (m, 3H), 8.15 (d, 1H), 8.00 (d, 1H), 7.90 (d, 1H), 7.50

(t, 2H), 4.40 (m, 1H), 3.30 (m, 3H), 2.90 (m, 4H), 1.80 (m, 4H); MS (EI) for $C_{23}H_{25}N_7O_2$: 432 (MH^+).

[1329] 3-amino-6-(3-[(3-fluorophenyl)acetyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1NMR (400 MHz, d_6 -DMSO): 10.80 (s, 1H), 9.10 (br s, 1H), 8.95 (br s, 1H), 8.80 (s, 1H), 8.70 (d, 1H), 8.40 (s, 1H), 7.80 (m, 2H), 7.40 (m, 2H), 7.30 (m, 2H), 7.10 (t, 1H), 4.30 (m, 1H), 3.75 (s, 2H), 3.22 (m, 3H), 2.80 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{24}H_{25}N_6O_2F$: 449 (MH^+).

[1330] 3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[(pyridin-3-ylacetyl)amino]phenyl}pyrazine-2-carboxamide: 1NMR (400 MHz, d_6 -DMSO): 11.15 (s, 1H), 9.28 (br s, 1H), 9.02 (m, 2H), 8.90 (d, 1H), 8.78 (s, 1H), 8.65 (d, 1H), 8.60 (d, 1H), 8.50 (s, 1H), 8.02 (m, 1H), 7.80 (m, 2H), 7.40 (t, 1H), 4.30 (m, 1H), 3.30 (m, 3H), 2.80 (m, 1H), 1.92 (m, 4H); MS (EI) for $C_{23}H_{25}N_7O_2$: 432 (MH^+).

[1331] 3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[(pyridin-3-ylcarbonyl)amino]phenyl}pyrazine-2-carboxamide: 1NMR (400 MHz, d_6 -DMSO): 11.30 (s, 1H), 9.80 (s, 1H), 9.30 (br s, 1H), 9.02 (m, 3H), 8.80 (m, 3H), 8.10 (d, 1H), 8.01 (m, 1H), 7.90 (d, 1H), 7.45 (t, 1H), 4.30 (m, 1H), 3.35 (m, 3H), 2.78 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{22}H_{23}N_7O_2$: 418 (MH^+).

[1332] 3-amino-6-(3-[(2,5-difluorophenyl)acetyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1NMR (400 MHz, d_6 -DMSO): 10.75 (s, 1H), 9.10 (br s, 1H), 9.01 (br s, 1H), 8.80 (s, 1H), 8.70 (d, 1H), 8.60 (s, 1H), 7.80 (m, 2H), 7.40 (t, 1H), 7.32 (m, 3H), 4.30 (m, 1H), 3.80 (s, 2H), 3.20 (m, 3H), 2.78 (m, 1H), 1.90 (m, 4H); MS (EI) for $C_{24}H_{24}N_6O_2F_2$: 467 (MH^+).

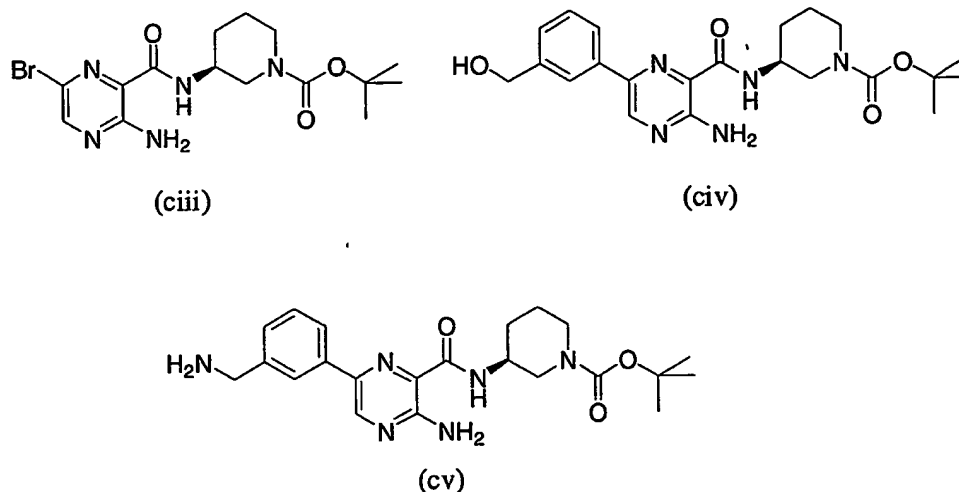
[1333] 3-amino-6-(3-[(4-fluorophenyl)acetyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 1NMR (400 MHz, d_6 -DMSO): 10.72 (s, 1H), 9.18 (br s, 1H), 9.02 (br s, 1H), 8.80 (s, 1H), 8.70 (d, 1H), 7.80 (m, 2H), 7.40 (m, 3H), 7.20 (m, 2H), 4.24 (m, 1H), 3.73 (s, 2H), 3.30 (m, 3H), 2.80 (m, 1H), 1.92 (m, 4H); MS (EI) for $C_{24}H_{25}N_6O_2F$: 449 (MH^+).

Example 67

[1334] Scheme 25 depicts three exemplary compounds, (ciii), (civ), and (cv), the latter two according to formula I, which are useful as intermediates to synthesize other compounds of the invention, particularly assembling -W-X-Y. The general methods

described above are used in various combinations to make compounds, (ciii), (civ), and (cv). Exemplary syntheses of such compounds follows

Scheme 25



1,1-dimethylethyl (3S)-3-[[[(3-amino-6-bromopyrazin-2-yl)carbonyl]amino]piperidine-1-carboxylate

[1335] 3-amino-6-bromopyrazine-2-carboxylic acid (3.63g, 16.65 mmol), HATU (9.49g, 25.0 mg) and DIEA (4.35 mL, 25.0 mmol) was added to a solution of (S)-3-amino-piperidine-1-carboxylic acid *tert*-butyl ester (4.00g, 19.97 mmol, commercially available from Astatech or Arch Corporation). It was stirred at room temperature overnight. The reaction mixture was poured into water (200 mL), and extracted with ethyl acetate (2 × 150 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography to give 1,1-dimethylethyl (3S)-3-[[[(3-amino-6-bromopyrazin-2-yl)carbonyl]amino}piperidine-1-carboxylate 5.00 g (75.0% yield): MS (EI) for C₁₅H₂₂BrN₅O₃: 401.12 (MH⁺).

1,1-dimethylethyl (3S)-3-[(3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2yl)carbonyl]
amino] piperidine-1-carboxylate

[1336] 1,1-dimethylethyl (3S)-3-([(3-amino-6-bromopyrazin-2-yl)carbonyl]amino) piperidine-1-carboxylate (765.0 mg, 1.91 mmol) was dissolved in DMF followed by addition of 3-hydroxymethyl phenylboronic acid (349.0 mg, 2.23 mmol), K₂CO₃ (660.0 mg, 4.77 mmol) and Pd(PPh₃)₄ (10 mol%, 220 mg). This mixture was stirred at 90 °C under nitrogen for 12 hours. The reaction mixture was poured into water (150 mL), and extracted with ethyl acetate (3 × 50 mL). The organic layers were washed with brine (50

mL) then dried over anhydrous sodium sulfate. Filtration, concentration and column chromatography on silica (50% ethylacetate/hexanes) gave a solid which was dried to afford 765.0 mg (92% yield) of 1,1-dimethylethyl (3S)-3-[(3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl)carbonyl) amino] piperidine-1-carboxylate: MS (EI) for $C_{22}H_{29}N_5O_4$: 428.23 (MH^+).

1,1-dimethylethyl (3S)-3-[(3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl)carbonyl) amino]piperidine-1-carboxylate

[1337] 1,1-Dimethylethyl (3S)-3-[(3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl)carbonyl)amino]piperidine-1-carboxylate (965.0 mg, 2.26 mmol) was dissolved in THF (20 mL) followed by the addition of diphenylphosphoryl azide (0.68 mL, 3.16 mmol) and DBU (0.50 mL, 3.16 mmol). It was stirred at room temperature overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2×50 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography (20% \rightarrow 50% Ethylacetate/hexanes) to give the corresponding azide, 588.0 mg (57.6 % yield) and then it was dissolved in the mixture of THF (20 mL) and H_2O (2.0 mL) followed by the addition of triphenylphosphine (1.02g, 3.88 mmol) and it was stirred at room temperature overnight. The reaction mixture was poured into water (100 mL), and extracted with ethyl acetate (2×50 mL) then dried over anhydrous sodium sulfate and filtration, concentration and purified by column chromatography (100% Ethylacetate \rightarrow 100% MeOH) to give 1,1-dimethylethyl (3S)-3-[(3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl)carbonyl) amino]piperidine-1-carboxylate, 435.0 mg (78.8 % yield): MS (EI) for $C_{22}H_{30}N_6O_3$: 427.24 (MH^+).

Example 68

3-Amino-6-[3-[(4-fluorophenyl)carbonyl]amino]methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

[1338] 1,1-dimethylethyl (3S)-3-[(3-amino-6-[3-(amino-methyl)phenyl]pyrazin-2-yl)carbonyl) amino]piperidine-1-carboxylate (150.0 mg, 0.35 mmol) and 4-fluorobenzoic acid (50.0 mg, 0.36 mmol) was dissolved in DMF (5mL) followed by the addition of HOBt (95.5 mg, 0.70 mmol) and EDCI (135.4 mg, 0.70 mmol). It was stirred at room temperature overnight. The reaction mixture was poured into water (20 mL), and extracted with ethyl acetate (2×30 mL) then dried over anhydrous sodium sulfate and

filtration, concentration and purified by column chromatography (50% Ethylacetate/hexanes) and subsequently it was treated with 4.0 M HCl in dioxane (20 mL) to give compound 3-Amino-6-[3-({[(4-fluorophenyl)carbonyl]amino}methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: 70.0 mg (46.6 % yield). ¹H NMR (400 MHz, d₄-MeOH): δ 8.62 (s, 1H), 8.16 (s, 1H), 7.98 (m, 3H), 7.48 (m, 2H), 7.20 (m, 3H), 4.67 (s, 2H), 4.38 (m, 1H), 3.45 (m, 1H), 3.38 (m, 1H), 3.18 (m, 1H), 3.00 (m, 1H), 2.18 (m, 2H), 1.80 (m, 2H); MS (EI) for C₂₄H₂₅N₆O₂F: 449.32 (MH⁺).

[1339] Using the same or analogous synthetic techniques and/or substituting with alternative reagents {respective corresponding amines, etc.}, the following compounds of the invention were prepared.

[1340] 3-amino-6-[3-({[(4-chlorophenyl)carbonyl]amino}methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ 9.50 (m, 2H), 9.30 (br, 1H), 8.78 (d, 1H), 8.20 (s, 1H), 8.00 (m, 3H), 7.60 (m, 2H), 7.40 (m, 2H), 5.00 (br, 2H), 4.67 (d, 2H), 4.38 (m, 1H), 3.20 (m, 3H), 2.80 (m, 1H), 1.80 (m, 4H); MS (EI) for C₂₄H₂₅N₆O₂Cl: 465.21 (MH⁺).

[1341] 3-amino-6-[3-({[(5-(4-nitrophenyl)furan-2-yl)carbonyl]amino}methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₄-MeOH): δ 8.62 (s, 1H), 8.38 (d, 2H), 8.10 (d, 3H), 7.98 (d, 1H), 7.50 (m, 2H), 7.30 (dd, 2H), 4.70 (s, 2H), 4.30 (br, 1H), 3.70 (m, 1H), 3.50 (m, 1H), 3.38 (m, 1H), 3.17 (m, 1H), 3.00 (m, 1H), 2.10 (m, 2H), 1.80 (m, 2H); MS (EI) for C₂₈H₂₈N₇O₅: 543.28 (MH⁺).

[1342] 3-amino-6-[3-({[(3-hydroxypyridin-2-yl)carbonyl]amino}methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₄-MeOH): δ; 8.70 (s, 1H), 8.40 (d, 2H), 8.10 (m, 1H), 8.00 (m, 1H), 7.40 (m, 1H), 7.30 (m, 1H), 4.80 (s, 1H), 4.40 (br, 1H), 4.08 (s, 2H), 3.70 (m, 1H), 3.50 (m, 1H), 3.40 (m, 1H), 3.20 (m, 1H), 3.10 (m, 1H), 2.10 (m, 2H), 1.80 (m, 2H); MS (EI) for C₂₃H₂₅N₇O₃: 448.28 (MH⁺).

[1343] 3-amino-N-[(3S)-piperidin-3-yl]-6-[3-({[(4-[(trifluoromethyl)oxy]phenyl)oxy]acetyl]amino}methyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ; 9.20 (m, 1H), 9.00 (m, 1H), 8.92 (m, 1H), 8.70 (d, 1H), 8.00 (m, 2H), 7.40 (m, 1H), 7.28 (m, 3H), 7.00 (m, 2H), 4.60 (s, 1H), 4.40 (d, 2H), 4.25 (m, 1H), 3.20 (m, 4H), 2.70 (m, 1H), 1.80 (m, 3H); MS (EI) for C₂₆H₂₇N₆O₄F₃: 545.26 (MH⁺).

[1344] 3-amino-6-[3-({[(5-methylisoxazol-3-yl)carbonyl]amino}methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d₆-DMSO): δ; 9.40 (m,

1H), 9.20 (m, 1H), 9.00 (m, 1H), 8.60 (d, 1H), 8.00 (m, 2H), 7.40 (m, 1H), 7.28 (m, 1H), 6.60 (s, 1H), 4.50 (d, 2H), 4.20 (m, 1H), 3.20 (m, 4H), 2.75 (m, 1H), 2.45 (s, 3H), 1.80 (m, 3H); MS (EI) for C₂₂H₂₅N₇O₃: 436.25 (MH⁺).

[1345] 3-amino-6-(3-([(isoxazol-5-ylcarbonyl)amino]methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d6-DMSO): δ; 9.70 (m, 1H), 9.20 (m, 1H), 9.00 (m, 1H), 8.70 (d, 1H), 8.68 (m, 1H), 8.15 (s, 1H), 8.00 (d, 1H), 7.40 (m, 1H), 7.30 (m, 1H), 7.20 (s, 1H), 4.50 (d, 2H), 4.20 (m, 1H), 3.20 (m, 4H), 2.75 (m, 1H), 1.80 (m, 3H); MS (EI) for C₂₁H₂₃N₇O₃: 422.23 (MH⁺).

[1346] 3-amino-6-[3-([(4-methyl-1,2,3-thiadiazol-5-yl)carbonyl]amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d6-DMSO): δ; 9.60 (m, 1H), 9.20 (m, 1H), 9.00 (m, 1H), 8.70 (d, 2H), 8.15 (s, 1H), 8.00 (d, 1H), 7.40 (m, 2H), 4.50 (d, 2H), 4.20 (m, 1H), 3.20 (m, 4H), 2.80 (s, 3H), 2.75 (m, 1H), 1.80 (m, 3H); MS (EI) for C₂₁H₂₄N₈O₂S: 453.25 (MH⁺).

[1347] 3-amino-6-[3-([(2,5-dichloro-3-thienyl)carbonyl]amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d6-DMSO): δ 9.40 (m, 1H), 9.20 (m, 1H), 9.80 (s, 1H), 8.70 (d, 1H), 8.20 (s, 1H), 8.00 (d, 1H), 7.60 (s, 1H), 7.40 (m, 2H), 4.50 (d, 2H), 4.30 (m, 1H), 3.70 (m, 1H), 3.50 (m, 1H), 3.20 (m, 2H), 2.80 (m, 1H), 1.80 (m, 3H); MS (EI) for C₂₂H₂₂N₆O₂SCl₂: 505.12 (M⁺).

[1348] 3-amino-6-[3-([(2,4-difluorophenyl)methyl]amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, d6-DMSO): δ 10.30 (br, 1H); 9.50 (m, 1H), 9.28 (m, 1H), 8.90 (m, 2H), 8.15 (m, 1H), 7.90 (m, 1H), 7.50 (m, 2H), 7.38 (m, 1H), 7.20 (m, 1H), 4.20 (m, 4H), 3.40 (m, 2H), 3.20 (m, 2H), 2.70 (m, 1H), 2.00 (m, 1H), 1.80 (m, 3H); MS (EI) for C₂₄H₂₆N₆OF₂: 453.25 (MH⁺).

Example 69

[1349] Example 69 describes synthesis of azepan derivatives according to formula I. Generally, but not necessarily, the synthetic strategies follow those described above.

3-Amino-N-azepan-3-yl-6-(3-([(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)pyrazine-2-carboxamide

[1350] To a solution of 3-amino-6-(3-([(1R)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)pyrazine-2-carboxylic acid (409 mg, 1.09 mmol) were added azepan-3-amine

(125 mg, 1.09 mmol), EDCI (252 mg, 1.31 mmol), HOBt (178 mg, 1.31 mmol) at ambient temperature. After being stirred for 2 h, the reaction mixture was quenched by addition of aqueous sodium bicarbonate followed by extraction with ethyl acetate (20 mL x 3). The aqueous phase was extracted once with additional ethyl acetate. The combined organic layers were washed with saturated aqueous sodium chloride then dried over anhydrous magnesium sulfate. Filtration and concentration in vacuo gave a crude oil, which was purified by preparative HPLC to yield the 3-amino-*N*-azepan-3-yl-6-(3-([(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl)phenyl)pyrazine-2-carboxamide (22 mg, 4.2 % yield): ¹H NMR (400 MHz, CD₃OD): δ 8.70 (m, 1H), 8.67 (m, 1H), 8.27 (m, 1H) 7.97 (m, 1H), 7.62 (t, 1H), 7.25 (m, 4H), 5.70 (m, 1H), 4.99 (m, 1H), 4.43 (m, 1H), 3.46 (m, 2H), 3.38 (m, 1H), 3.15 (m, 2H), 2.94 (m, 1H), 2.61 (m, 1H), 2.17 (m, 2H), 1.95 (m, 4H), 1.62 (m, 1H); MS (EI) for C₂₇H₃₀N₆O₂: 471.32 (MH⁺).

3-Amino-6-[3-([(2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino} carbonyl)phenyl]-
N-[(3*S*)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide

[1351] 3-Amino-6-[3-([(1*S*,2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino}carbonyl)phenyl] pyrazine-2-carboxylic acid (1.0 eq) was dissolved in DMF-THF and treated with (3*S*)-1-(phenylmethyl)azepan-3-amine (1.46 eq), EDCI (1.2 eq) and HOBt (1.2 eq) at ambient temperature. After being stirred overnight, the reaction mixture was quenched by addition of aqueous sodium bicarbonate followed by extraction with ethyl acetate (20 mL x 3). The aqueous phase was extracted once with additional ethyl acetate. The combined organic layers were washed with saturated aqueous sodium chloride then dried over anhydrous magnesium sulfate. Filtration and concentration in vacuo gave a crude oil, which was purified by preparative HPLC to yield a crude oil, which was dissolved in ethyl acetate (10mL), and treated with 1M HCl in ether (5mL). The resulting precipitation was filtered, washed with methanol (2 x 20mL), and dried to afford 122 mg (12% yield) of 3-amino-6-[3-([(2*S*)-2-hydroxy-2,3-dihydro-1*H*-inden-1-yl]amino}carbonyl)phenyl]-N-[(3*S*)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide (122 mg, 15 % yield). : ¹H NMR (400 MHz, DMSO-*d*₆): δ 8.98 (s, 1H), 8.90 (m, 1H), 8.75 (s, 1H), 8.63 (m, 2H), 8.11 (m, 1H), 7.95 (m, 2H), 7.62 (m, 2H), 7.58 (m, 1H), 7.42 (m, 1H), 7.20 (m, 3H), 7.08 (m, 1H), 5.50 (m, 1H), 4.51 (m, 1H), 4.49 (m, 1H), 4.46 (m, 1H), 4.40 (m, 1H), 3.50 (m, 1H), 3.65 (m, 2H), 3.20 (m, 2H), 2.85 (m, 1H), 2.10 (m, 1H), 1.80 (m, 2H), 1.58 (m, 1H) ; MS (EI) for C₃₄H₃₆N₆O₃: 577.08 (MH⁺).

- [1352] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino}carbonyl)phenyl]pyrazine-2-carboxamide was prepared using the same synthetic procedure as above followed by hydrogenation (1 atm) with 10% palladium hydroxide in MeOH: ^1H NMR (400 MHz, DMSO- d_6): δ 9.18 (bs, 1H), 8.97 (m, 2H), 8.82 (m, 1H), 8.61 (m, 2H), 8.33 (m, 1H), 7.92 (m, 1H), 7.70 (bs, 2H), 7.59 (t, 1H), 7.27 (m, 2H), 7.20 (m, 1H), 5.50 (m, 1H), 4.53 (m, 1H), 4.35 (m, 1H), 3.27 (m, 1H), 3.09 (m, 2H), 1.98 (m, 1H), 1.84 (m, 4H), 1.57 (m, 2H); MS (EI) for $\text{C}_{27}\text{H}_{30}\text{N}_6\text{O}_3$: 487.07 (MH^+).
- [1353] 1,1-dimethylethyl (3S)-3-{{[(3-amino-6-bromopyrazin-2-yl)carbonyl]amino}azepane-1-carboxylate: (3S)-3-aminoazepan-2-one is prepared by either the resolution of racemic α -amino- ϵ -caprolactam (Sigma-Aldrich) by the method of E. M. Rezler, R. R. Fenton, W. J. Esdale, M. J. McKeage, P. J. Russell, and T. W. Hambley *J. Med. Chem.* 1997, 40, 3508 – 3515, or by chiral synthesis from L-lysine using the method of R. Pellegata, M. Pinza, and G. Pifferi, *Synthesis*, 1978, 614 – 616. (3S)-3-[(Triphenylmethyl)amino]azepan-2-one and (3S)-N-(triphenylmethyl)azepan-3-amine were prepared using similar methods.
- [1354] 1,1-dimethylethyl (3S)-3-[(triphenylmethyl)amino]azepane-1-carboxylate: To an ice-cooled, stirred solution of (3S)-N-(triphenylmethyl)azepan-3-amine (30.0 mmol) in 60 mL of THF, was added 9.85 g (45.1 mmol, 1.5 eq.) of di-*tert*-butyl dicarbonate. The ice bath was removed, and the reaction stirred at room temperature for 19 h and concentrated. The crude oil was purified by silica gel chromatography (55 mm column, 7.5 inches of silica gel) using 10% EtOAc in hexanes as eluent. Concentration afforded 13.7 g (100% yield) of product as a solid.
- [1355] 1,1-dimethylethyl (3S)-3-aminoazepane-1-carboxylate: To 13.7 g (30 mmol) of 1,1-dimethylethyl (3S)-3-[(triphenylmethyl)amino]azepane-1-carboxylate was carefully added 100 mL MeOH, 75 mL EtOH, and 1.00 g of 10% Palladium on charcoal. The mixture was hydrogenated on a Parr shaker under 30 psi of hydrogen gas for 15 h. The mixture was filtered through a pad of celite, and the filtrate concentrated to afford a quantitative yield (6.43 g) of product. The product was used as such without further purification.
- [1356] 1,1-dimethylethyl (3S)-3-{{[(3-amino-6-bromopyrazin-2-yl)carbonyl]amino}azepane-1-carboxylate: A flask was charged with 6.43 g (30 mmol) of 1,1-dimethylethyl (3S)-3-aminoazepane-1-carboxylate, 109 mL THF, 5.49 g of 3-amino-6-bromopyrazine-2-carboxylic acid (25.2 mmol, 0.84 eq.), 5.00 mL TEA (35.9 mmol, 1.20 eq.), 5.11 g

(37.8 mmol, 1.26 eq.) of 1-hydroxybenzotriazole (HOBT), and 7.25 g (37.8 mmol, 1.26 eq.) of 1-[3-(dimethylamino)propyl]-3-ethylcarbodiimide hydrochloride (EDCI). The reaction was allowed to stir for 16 h. The reaction was diluted with EtOAc, washed with 2 x sat. aqueous NaHCO₃, 1 x sat. aqueous NaCl, dried (Na₂SO₄), filtered, and concentrated. The crude material was purified by silica gel chromatography on a 55 mm column (7.5 inches of silica gel) using 30 – 50% EtOAc in hexanes as eluent. Concentration afforded 4.57 g (43.8% yield) of product as a solid.

[1357] The 1,1-dimethylethyl (3S)-3-[[3-(3-amino-6-bromopyrazin-2-yl)carbonyl]amino]azepane-1-carboxylate could be used in a manner analogous to that used for 1,1-dimethylethyl (3S)-3-[[3-(3-amino-6-bromopyrazin-2-yl)carbonyl]amino]piperidine-1-carboxylate to prepare compounds that are part of the invention.

[1358] Using the same or analogous synthetic techniques and/or substituting with alternative reagents, the following azepan compounds of the invention were prepared:

[1359] 3-Amino-6-{3-[(2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl}-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.20 (d, 1H), 8.95 (m, 3H), 8.85 (s, 1H), 8.50 (d, 1H), 7.97 (m, 1H), 7.50 (m, 3H), 7.41 (m, 1H), 7.32 (m, 4H), 7.05 (m, 1H), 5.60 (m, 1H), 4.35 (m, 1H), 4.29 (m, 1H), 4.20 (m, 1H), 3.08 (m, 2H), 2.91 (m, 2H), 2.45 (m, 1H), 2.31 (m, 1H), 1.80 (m, 3H), 1.42 (m, 1H); MS (EI) for C₃₄H₃₆N₆O₂: 461.07 (MH⁺).

[1360] 3-Amino-N-[(3S)-azepan-3-yl]-6-{3-[(2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl}pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.64 (bs, 1H), 9.32 (bs, 1H), 9.21 (d, 1H), 8.98 (m, 2H), 8.70 (bs, 1H), 8.39 (d, 1H), 7.93 (d, 1H), 7.58 (t, 1H), 7.30 (m, 4H), 5.62 (m, 1H), 4.35 (m, 1H), 3.21 (m, 2H), 3.18 (m, 1H), 3.02 (m, 2H), 2.85 (m, 1H), 2.43 (m, 1H), 2.18 (m, 1H), 1.93 (m, 1H), 1.80 (m, 4H), 1.55 (m, 1H); MS (EI) for C₂₇H₃₀N₆O₂: 471.10 (MH⁺).

[1361] 3-Amino-N-[(3S)-azepan-3-yl]-6-(3-methylphenyl)pyrazine-2-carboxamide: ¹H NMR (400 MHz, CD₃OD): δ 8.60 (s, 1H), 7.88 (m, 2H), 7.37 (m, 1H), 7.23 (m, 1H), 4.45 (m, 1H), 3.40 (m, 4H), 2.43 (s, 3H), 2.21 (m, 1H), 2.00 (m, 2H), 1.65 (m, 1H), 1.32 (m, 1H), 0.98 (m, 1H); MS (EI) for C₁₈H₂₃N₅O: 326.28 (MH⁺).

[1362] 3-Amino-N-[(3S)-azepan-3-yl]-6-{3-[[4-chloro-2-fluorophenyl)methyl]amino}carbonyl]phenyl}pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.50 (m, 2H), 9.15 (bs, 1H), 9.02 (d, 1H), 8.92 (s, 1H), 8.73 (s, 1H),

8.31 (d, 1H), 7.86 (d, 1H), 7.56 (t, 1H), 7.45 (m, 2H), 7.25 (m, 1H), 4.94 (bs, 2H), 4.49 (m, 2H), 4.40 (m, 1H), 3.33 (m, 2H), 3.23 (m, 1H), 3.00 (m, 1H), 2.08 (m, 1H), 1.80 (m, 3H), 1.55 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2FCl$: 497.03 (MH^+).

[1363] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(2,3-difluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): δ 9.30 (t, 1H), 8.98 (m, 2H), 8.70 (m, 2H), 8.59 (s, 1H), 8.32 (s, 1H), 7.89 (d, 1H), 7.72 (bs, 1H), 7.58 (t, 1H), 7.34 (m, 1H), 7.20 (m, 2H), 4.60 (m, 2H), 4.36 (m, 1H), 3.35 (m, 1H), 3.22 (m, 1H), 3.09 (m, 1H), 2.67 (m, 1H), 2.33 (m, 1H), 1.99 (m, 1H), 1.85 (m, 3H), 1.58 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2F_2$: 481.08 (MH^+).

[1364] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(3,4-difluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): δ 9.48 (m, 1H), 9.37 (m, 1H), 8.99 (m, 2H), 8.95 (s, 1H), 8.32 (d, 1H), 7.85 (d, 1H), 7.55 (t, 1H), 7.43 (m, 2H), 7.23 (m, 1H), 4.50 (m, 3H), 3.34 (m, 2H), 3.24 (m, 2H), 3.07 (m, 1H), 2.05 (m, 1H), 1.80 (m, 4H), 1.58 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2F_2$: 481.08 (MH^+).

[1365] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(3,5-difluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): δ 9.46 (t, 2H), 9.18 (bs, 1H), 8.95 (m, 3H), 8.68 (s, 1H), 8.33 (m, 1H), 7.91 (d, 1H), 7.70 (bs, 1H), 7.60 (t, 1H), 7.12 (m, 3H), 4.54 (m, 2H), 4.39 (m, 1H), 3.38 (m, 2H), 3.24 (m, 1H), 3.06 (m, 1H), 2.00 (m, 1H), 1.89 (m, 4H), 1.57 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2F_2$: 481.07 (MH^+).

[1366] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(3-chloro-2,6-difluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): δ 9.58 (bs, 1H), 9.30 (m, 1H), 9.23 (bs, 1H), 9.00 (d, 1H), 8.93 (s, 1H), 8.63 (s, 1H), 8.52 (bs, 2H), 8.33 (m, 1H), 7.86 (m, 1H), 7.79 (m, 1H), 7.60 (m, 2H), 7.32 (t, 1H), 7.18 (t, 1H), 4.58 (m, 2H), 4.41 (m, 1H), 4.06 (m, 2H), 3.31 (m, 1H), 3.22 (m, 2H), 3.01 (m, 1H), 2.00 (m, 1H), 1.90 (m, 2H), 1.80 (m, 2H), 1.59 (m, 1H); MS (EI) for $C_{25}H_{25}N_6O_2F_2Cl$: 515.03 (MH^+).

[1367] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(2,3-dihydro-1,4-benzodioxin-6-yl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): δ 9.48 (bs, 1H), 9.38 (m, 1H), 9.14 (bs, 1H), 9.10 (m, 1H), 8.95 (s, 1H), 8.69 (s, 1H), 8.32 (m, 1H), 7.89 (m, 1H), 7.57 (m, 1H), 6.83 (s, 3H), 4.41 (m, 2H), 3.45 (m, 4H), 3.17

(m, 1H), 2.00 (m, 1H), 1.89 (m, 3H), 1.59 (m, 1H); MS (EI) for $C_{27}H_{30}N_6O_4$: 503.10. (MH⁺)

[1368] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(2-chloro-6-fluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.34 (bs, 1H), 9.09 (m, 1H), 8.97 (d, 1H), 8.92 (s, 1H), 8.60 (s, 2H), 8.55 (bs, 2H), 8.34 (m, 1H), 7.85 (m, 1H), 7.54 (m, 2H), 7.44 (m, 1H), 7.39 (m, 2H), 7.25 (m, 1H), 4.63 (m, 2H), 4.40 (m, 1H), 3.25 (m, 2H), 3.03 (m, 1H), 2.00 (m, 1H), 1.90 (m, 1H), 1.83 (m, 2H), 1.59 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2FCl$: 497.31 (MH⁺).

[1369] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(2,6-difluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.20 (bs, 1H), 9.05 (m, 1H), 8.97 (bs, 1H), 8.90 (s, 1H), 8.82 (m, 1H), 8.51 (s, 1H), 8.30 (m, 1H), 7.83 (m, 1H), 7.69 (bs, 1H), 7.53 (t, 1H), 7.36 (m, 1H), 7.10 (m, 2H), 4.55 (m, 2H), 4.34 (m, 1H), 3.25 (m, 3H), 3.06 (m, 2H), 1.90 (m, 1H), 1.83 (m, 3H), 1.57 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2F_2$: 481.29 (MH⁺).

[1370] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(3-chloro-4-fluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.46 (m, 1H), 9.27 (bs, 1H), 8.94 (m, 2H), 8.92 (s, 1H), 8.67 (m, 1H), 8.30 (d, 1H), 7.70 (bs, 1H), 7.60 (m, 2H), 7.56 (m, 2H), 7.36 (m, 2H), 4.50 (m, 3H), 3.32 (m, 2H), 3.22 (m, 2H), 3.03 (m, 1H), 2.01 (m, 1H), 1.85 (m, 3H), 1.59 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2FCl$: 497.25 (MH⁺).

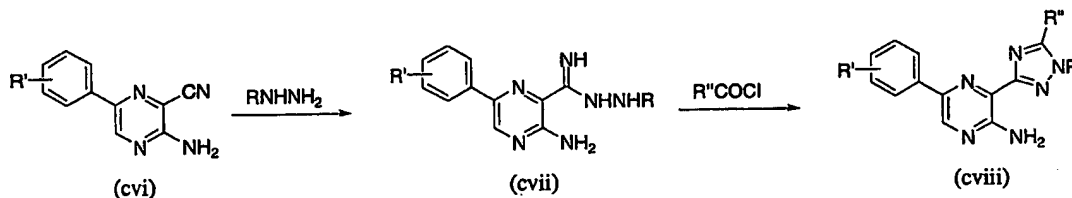
[1371] 3-Amino-N-[(3S)-azepan-3-yl]-6-[3-({[(3-chloro-2-fluorophenyl)methyl]amino}carbonyl)phenyl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, DMSO-d₆): δ 9.36 (m, 1H), 9.14 (m, 1H), 8.92 (s, 1H), 8.87 (m, 2H), 8.62 (m, 1H), 8.10 (m, 1H), 7.88 (m, 2H), 7.57 (t, 1H), 7.48 (m, 1H), 7.39 (m, 1H), 7.19 (m, 1H), 4.57 (m, 1H), 4.36 (bs, 1H), 3.23 (m, 1H), 3.04 (m, 1H), 1.99 (m, 1H), 1.90 (m, 3H), 1.79 (m, 1H), 1.57 (m, 1H); MS (EI) for $C_{25}H_{26}N_6O_2FCl$: 497.04 (MH⁺).

Example 70

[1372] Scheme 26 depicts another strategy for making exemplary compounds according to formula I where A is a triazole. Nitrile intermediates (cvi) were converted to aminoamidines (cvii), which were acylated and the intermediates cyclized to form

triazoles (cviii). Intermediate (cvi) can also be used to make other heterocycles for A, according to formula I.

Scheme 26



[1373] 3-{5-Amino-6-[[2-(1,3-benzothiazol-2-yl)hydrazino](imino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide: A mixture of 3-(5-amino-6-cyanopyrazin-2-yl)-N-(phenylmethyl)benzamide (100 mg, 0.3 mmol) and 2-hydrazinobenzothiazole (99 mg, 0.6 mmol) in EtOH (3 mL) was heated at 100-110 °C in a pressure vessel for 24 h. The mixture was then cooled, and the solid filtered and washed several times with ether to give the 3-{5-amino-6-[[2-(1,3-benzothiazol-2-yl)hydrazino](imino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide as a yellow solid (126 mg, 85% yield). ¹H NMR (400 MHz, *d*₆-DMSO): δ 11.50 (s, 1H), 9.27-9.16 (m, 1H), 8.81 (s, 1H), 8.58 (s, 1H), 8.37-8.29 (m, 1H), 8.0-7.45 (m, 4H), 7.45-6.10 (m, 11H), 4.61-4.47 (m, 2H). MS (EI) for C₂₆H₂₂N₈OS: 495 (MH⁺).

[1374] 3-[5-Amino-6-(5-phenyl-1-pyridin-2-yl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide: To a solution of 3-{5-amino-6-[imino(2-pyridin-2-yl)hydrazino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide (50 mg, 0.11 mmol) in dry pyridine (1 mL) at room temperature was added dropwise benzoyl chloride (0.02 mL, 0.17 mmol), and the resulting mixture was then heated at 120 °C for 14 h in a pressure vessel. The pyridine was removed under reduced pressure and the residue was purified by preparative HPLC to give 3-[5-amino-6-(5-phenyl-1-pyridin-2-yl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide as a yellow solid (41 mg, 69% yield). ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.86 (s, 1H), 8.55 (s, 1H), 8.54-8.50 (m, 1H), 8.26-8.21 (m, 1H), 8.20-8.13 (m, 1H), 7.95-7.87 (m, 2H), 7.67-7.42 (m, 9H), 7.39-7.30 (m, 5H), 7.28-7.20 (m, 1H), 4.55-4.50 (m, 2H). MS (EI) for C₃₁H₂₄N₈O: 525 (MH⁺).

[1375] Using the same or similar synthetic techniques, substituting with the appropriate reagents such as the respective amines, the following compounds of the invention were prepared:

- [1376] 3-{5-Amino-6-[imino(2-methylhydrazino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.15 (m, 1H), 8.95-8.91 (m, 1H), 8.54 (s, 1H), 8.31 (bs, 1H), 7.93-7.88 (m, 1H), 7.93-7.42 (bs, 2H), 7.62-7.55 (m, 1H), 7.38-7.31 (m, 4H), 7.29-7.23 (m, 1H), 4.57-4.51 (m, 1H), 2.89 (bs, 3H); MS (EI) for $\text{C}_{20}\text{H}_{21}\text{N}_7\text{O}$: 376.16 (MH^+).
- [1377] 3-{5-Amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.15 (m, 1H), 8.74 (bs, 2H), 8.55-8.51 (m, 1H), 8.31-8.25 (m, 1H), 7.89-7.84 (m, 1H), 7.59-7.52 (m, 1H), 7.37-7.20 (m, 6H), 6.98-6.92 (m, 2H), 6.80-6.71 (m, 1H), 4.57-4.50 (m, 2H); MS (EI) for $\text{C}_{25}\text{H}_{23}\text{N}_7\text{O}$: 438.15 (MH^+).
- [1378] 3-{5-Amino-6-[imino(morpholin-4-ylamino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.14 (m, 1H), 8.90 (bs, 1H), 8.75 (s, 1H), 8.53-8.49 (m, 1H), 8.28-8.23 (m, 1H), 7.88-7.83 (m, 1H), 7.58-7.51 (m, 1H), 7.50-7.00 (m, 6H), 6.65 (bs, 2H), 4.56-4.49 (m, 2H), 3.80-3.71 (m, 4H), 2.80-2.65 (m, 4H); MS (EI) for $\text{C}_{23}\text{H}_{25}\text{N}_7\text{O}_2$: 432.15 (MH^+).
- [1379] 3-(5-Amino-6-{imino[(4-methylpiperazin-1-yl)amino]methyl}pyrazin-2-yl)-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.80 (bs, 2H), 9.20-9.12 (m, 1H), 8.86 (bs, 1H), 8.50 (s, 1H), 8.22 (bs, 1H), 7.91-7.86 (m, 1H), 7.60-7.53 (m, 1H), 7.36-7.20 (m, 5H), 4.56-4.48 (m, 2H), 3.40-3.15 (m, 4H), 2.92-2.75 (m, 7H); MS (EI) for $\text{C}_{24}\text{H}_{28}\text{N}_8\text{O}$: 445.18 (MH^+).
- [1380] 3-{5-Amino-6-[imino(piperidin-1-ylamino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.14 (m, 1H), 8.98 (bs, 1H), 8.73 (s, 1H), 8.53-8.49 (m, 1H), 8.27-8.22 (m, 1H), 7.88-7.83 (m, 1H), 7.58-7.51 (m, 1H), 7.38-7.18 (m, 5H), 6.51 (bs, 2H), 4.56-4.49 (m, 2H), 2.74-2.60 (m, 4H), 1.74-1.62 (m, 4H), 1.50-1.38 (m, 2H); MS (EI) for $\text{C}_{24}\text{H}_{27}\text{N}_7\text{O}$: 430.18 (MH^+).
- [1381] 3-{5-Amino-6-[(azepan-1-ylamino)(imino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -DMSO): δ 11.23 (bs, 1H), 9.80 (bs, 1H), 9.17-9.08 (m, 2H), 8.92 (s, 1H), 8.49-8.43 (m, 1H), 8.16-8.08 (m, 1H), 7.92-7.86 (m, 1H), 7.60-7.53 (m, 1H), 7.38-7.05 (m, 6H), 4.57-4.47 (m, 2H), 3.15-2.95 (m, 4H), 1.85-1.55 (m, 8H); MS (EI) for $\text{C}_{25}\text{H}_{29}\text{N}_7\text{O}$: 444.20 (MH^+).
- [1382] 3-{5-Amino-6-[imino({(2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl}amino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ^1H NMR (400 MHz, d_6 -

DMSO): δ 9.79 (bs, 1H), 9.37 (bs, 1H), 9.16-9.10 (m, 1H), 8.95 (s, 1H), 8.50-8.46 (m, 1H), 8.17-8.10 (m, 1H), 7.92-7.87 (m, 1H), 7.61-7.54 (m, 1H), 7.38-7.18 (m, 5H), 7.04 (bs, 2H), 4.55-4.48 (m, 2H), 3.55-3.48 (m, 2H), 3.42-3.32 (m, 1H), 3.28 (s, 3H), 3.27-3.18 (m, 1H), 2.85-2.74 (m, 1H), 2.10-1.72 (m, 4H), 1.54-1.38 (m, 1H); MS (EI) for $C_{25}H_{29}N_7O_2$: 460.18 (MH^+).

[1383] 3-[5-Amino-6-(imino{2-[4-(trifluoromethyl)pyrimidin-2-yl]hydrazino}methyl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.25-9.05 (m, 2H), 8.99 (s, 1H), 8.79 (bs, 1H), 8.47-8.43 (m, 1H), 8.14-8.08 (m, 1H), 7.95-7.90 (m, 1H), 7.62-7.41 (m, 4H), 7.39-7.21 (m, 6H), 6.91 (bs, 1H), 4.55-4.50 (m, 2H); MS (EI) for $C_{24}H_{20}N_9OF_3$: 508.09 (MH^+).

[1384] 3-[5-Amino-6-(1,5-diphenyl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.20-9.13 (m, 1H), 8.84 (s, 1H), 8.56-8.52 (m, 1H), 8.25-8.20 (m, 1H), 7.91-7.85 (m, 1H), 7.60-7.43 (m, 12H), 7.38-7.30 (m, 5H), 7.28-7.22 (m, 1H), 4.55-4.50 (m, 2H); MS (EI) for $C_{32}H_{25}N_7O$: 524.12 (MH^+).

[1385] 3-[5-Amino-6-(1,5-dimethyl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.21-9.14 (m, 1H), 8.75 (s, 1H), 8.54-8.50 (m, 1H), 8.22-8.17 (m, 1H), 7.90-7.85 (m, 1H), 7.62-7.55 (m, 2H), 7.40-7.20 (m, 6H), 4.57-4.50 (m, 2H), 3.94 (s, 3H), 2.55 (s, 3H); MS (EI) for $C_{22}H_{21}N_7O$: 400.16 (MH^+).

[1386] 3-[5-Amino-6-(1-methyl-5-phenyl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.22-9.13 (m, 1H), 8.79 (s, 1H), 8.56-8.52 (m, 1H), 8.24-8.19 (m, 1H), 7.95-7.86 (m, 3H), 7.68-7.51 (m, 5H), 7.40-7.21 (m, 6H), 4.57-4.48 (m, 2H), 4.13 (s, 3H); MS (EI) for $C_{27}H_{23}N_7O$: 462.11 (MH^+).

[1387] 3-{5-Amino-6-[imino(2-pyridin-2-ylhydrazino)methyl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.46 (bs, 1H), 9.26-9.19 (m, 1H), 8.74 (s, 1H), 8.59-8.55 (m, 1H), 8.36-8.30 (m, 1H), 8.16-8.12 (m, 1H), 7.91-7.86 (m, 1H), 7.70-7.62 (m, 1H), 7.60-7.53 (m, 1H), 7.40-7.32 (m, 4H), 7.30-7.22 (m, 1H), 6.90-6.82 (m, 1H), 6.79-6.70 (m, 3H), 4.59-4.53 (m, 2H); MS (EI) for $C_{24}H_{22}N_8O$: 439.15 (MH^+).

- [1388] 3-[5-Amino-6-(1-methyl-5-piperidin-2-yl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.42-9.23 (m, 2H), 9.21-9.13 (m, 1H), 8.79 (s, 1H), 8.52-8.48 (m, 1H), 8.20-8.14 (m, 1H), 7.92-7.86 (m, 1H), 7.68-7.54 (m, 2H), 7.39-7.31 (m, 4H), 7.30-7.22 (m, 1H), 4.92-4.81 (m, 1H), 4.57-4.49 (m, 2H), 4.06 (s, 3H), 3.22-3.02 (m, 2H), 2.23-2.14 (m, 1H), 1.95-1.60 (m, 5H); MS (EI) for C₂₆H₂₈N₈O: 469.17 (MH⁺).
- [1389] 3-[5-Amino-6-(5-methyl-1-pyridin-2-yl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.22-9.15 (m, 1H), 8.82 (s, 1H), 8.66-8.61 (m, 1H), 8.56-8.53 (m, 1H), 8.25-8.20 (m, 1H), 8.17-8.11 (m, 1H), 8.10-8.05 (m, 1H), 7.92-7.87 (m, 1H), 7.63-7.52 (m, 4H), 7.39-7.31 (m, 4H), 7.29-7.22 (m, 1H), 4.57-4.50 (m, 2H), 2.90 (s, 3H); MS (EI) for C₂₆H₂₂N₈O: 463.15 (MH⁺).
- [1390] 3-{5-Amino-6-[(2-ethylhydrazino)(imino)methyl]pyrazin-2-yl}-*N*-(2,3-dihydro-1*H*-inden-1-yl)benzamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.8 (s, 1H), 8.7 (s, 1H), 8.2 (m, 1H), 7.80 (m, 1H), 7.4 (m, 2H), 7.2 (m, 3H), 5.8 (m, 1H), 2.8 (m, 2H), 2.4 (m, 2H), 2.0 (m, 2H), 1.5 (m, 3H). MS (EI) for C₂₃H₂₅N₇O: 416 (MH⁺).
- [1391] 3-{5-Amino-6-[imino(2-methylhydrazino)methyl]pyrazin-2-yl}-*N*-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.96-8.91 (m, 1H), 8.64 (s, 1H), 8.53-8.50 (m, 1H), 8.30-8.24 (m, 1H), 7.92-7.87 (m, 1H), 7.58-7.52 (m, 1H), 7.32-7.17 (m, 4H), 6.03 (s, 2H), 5.67-5.58 (m, 1H), 5.26 (s, 1H), 3.08-2.96 (m, 1H), 2.94-2.85 (m, 1H), 2.84 (s, 3H), 2.50-2.44 (m, 1H), 2.07-1.99 (m, 1H). MS (EI) for C₂₂H₂₃N₇O: 402 (MH⁺).
- [1392] 3-{5-Amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl}-*N*-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.00-8.91 (m, 1H), 8.78-8.68 (m, 1H), 8.57-8.53 (m, 1H), 8.36-8.30 (m, 1H), 7.95-7.89 (m, 1H), 7.61-7.52 (m, 1H), 7.33-7.14 (m, 5H), 6.99-6.93 (m, 2H), 6.78-6.71 (m, 1H), 6.50 (s, 2H), 5.68-5.57 (m, 1H), 3.08-2.96 (m, 1H), 2.95-2.82 (m, 1H), 2.56-2.46 (m, 1H), 2.10-1.97 (m, 1H). MS (EI) for C₂₇H₂₅N₇O: 464 (MH⁺).
- [1393] Phenylmethyl 3-{3-[3-amino-6-(3-{[(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl}phenyl)pyrazin-2-yl]-1-methyl-1*H*-1,2,4-triazol-5-yl}piperidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.93-8.88 (m, 1H), 8.74 (s, 1H), 8.53-8.49 (m, 1H), 8.20-8.14 (m, 1H), 7.93-7.85 (m, 1H), 7.61-7.54 (m, 1H), 7.42-7.15 (m, 11H), 5.67-5.57

- (m, 1H), 5.10 (s, 2H), 4.18-3.88 (m, 6H), 2.12-1.93 (m, 2H), 1.88-1.70 (m, 2H), 1.64-1.42 (m, 2H). MS (EI) for $C_{36}H_{36}N_8O_3$: 629 (MH^+).
- [1394] Phenylmethyl 3-{3-[3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate: 1H NMR (400 MHz, d_4 -MeOH): δ 8.8 (s, 1H), 8.7 (s, 1H), 8.2 (m, 1H), 7.80 (m, 1H), 7.4 (m, 2H), 7.3 (m, 5H), 7.2 (m, 3H), 5.8 (m, 1H), 2.8 (m, 2H), 2.4 (m, 2H), 2.2-3.2 (m, 7H), 1.8-2.0 (m, 9H). MS (EI) for $C_{37}H_{38}N_8O_3$: 643 (MH^+).
- [1395] 3-Amino-6-bromo-*N'*-pyridin-2-ylpyrazine-2-carboximidohydrazide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.00 (s, 1H), 8.95 (m, 1H), 8.55 (s, 1H), 8.10 (d, 1H), 7.95 (d, 1H), 7.50 (m, 3H), 7.30-7.10 (m, 4H), 5.60 (m, 2H), 3.0-2.8 (m, 2H), 2.40 (m, 1H), 2.00 (m, 1H). MS (EI) for $C_{10}H_{10}BrN_7$: 356 (MH^+).
- [1396] 3-[5-Amino-6-(1-ethyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-(2,3-dihydro-1H-inden-1-yl)benzamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.8 (s, 1H), 8.7 (s, 1H), 8.2 (m, 1H), 7.80 (m, 1H), 7.4 (m, 2H), 7.2 (m, 3H), 5.8 (m, 1H), 2.8 (m, 2H), 2.4 (m, 2H), 2.2-3.2 (m, 5H), 1.8-2.0 (m, 9H). MS (EI) for $C_{29}H_{32}N_8O$: 509 (MH^+).
- [1397] 3-[5-Amino-6-(1-methyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-[[1S)-2,3-dihydro-1H-inden-1-yl]benzamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.57-8.49 (m, 2H), 8.12-8.05 (m, 1H), 7.89-7.82 (m, 1H), 7.54-7.46 (m, 1H), 7.28-7.06 (m, 4H), 5.67-5.56 (m, 1H), 3.96 (s, 3H), 3.57-3.24 (m, 3H), 3.10-2.87 (m, 3H), 2.60-2.46 (m, 1H), 2.15-1.40 (m, 6H). MS (EI) for $C_{28}H_{30}N_8O$: 495 (MH^+).
- [1398] 3-{5-Amino-6-[imino(2-pyridin-4-ylhydrazino)methyl]pyrazin-2-yl}-*N*-(2,3-dihydro-1H-inden-1-yl)benzamide: 1H NMR (400 MHz, d_6 -DMSO): δ 10.9 (s, 1H), 8.95 (m, 2H), 8.55 (s, 1H), 8.30-8.10 (m, 3H), 7.95-7.00 (m, 10H), 5.60 (m, 2H), 3.0-2.8 (m, 2H), 2.40 (m, 1H), 2.00 (m, 1H). MS (EI) for $C_{26}H_{24}N_8O$: 465 (MH^+).
- [1399] Phenylmethyl 3-[3-(3-amino-6-bromopyrazin-2-yl)-1-pyridin-2-yl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate: 1H NMR (400 MHz, d_4 -MeOH): δ 8.20 (s, 1H), 7.40-7.20 (m, 9H), 5.10 (m, 2H), 4.00-3.60 (m, 4H), 3.2 (m, 1H), 2.0-1.40 (m, 4H). MS (EI) for $C_{24}H_{23}BrN_8O_8$: 536 (MH^+).
- [1400] Phenylmethyl (3R)-3-{3-[3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate: 1H NMR (400 MHz, d_4 -MeOH): δ 8.8 (s, 1H), 8.7 (s, 1H), 8.2 (m, 1H),

7.80 (m, 1H), 7.4 (m, 2H), 7.3 (m, 5H), 7.2 (m, 3H), 5.8 (m, 1H), 2.8 (m, 2H), 2.4 (m, 2H), 2.2-3.2 (m, 7H), 1.8-2.0 (m, 9H). MS (EI) for C₃₇H₃₈N₈O₃: 643 (MH⁺).

[1401] 3-(5-Amino-6-{1-ethyl-5-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide: ¹H NMR (400 MHz, d₄-MeOH): δ 8.8 (s, 1H), 8.7 (s, 1H), 8.2 (m, 1H), 7.80 (m, 1H), 7.4 (m, 2H), 7.2 (m, 3H), 5.8 (m, 1H), 2.8 (m, 2H), 2.4 (m, 2H), 2.2-3.2 (m, 5H), 1.8-2.0 (m, 9H). MS (EI) for C₂₉H₃₂N₈O: 509 (MH⁺).

[1402] Phenylmethyl (3R)-3-{3-[3-amino-6-(3-{[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl}phenyl)pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl}piperidine-1-carboxylate ¹H NMR (400 MHz, d₄-MeOH): δ 8.95 (m, 1H), 8.60 (m, 1H), 8.20 (d, 1H), 7.95 (d, 1H), 7.60 (m, 1H), 7.40-7.20 (m, 9H), 5.65 (m, 1H), 5.20 (m, 2H), 4.2-3.8 (m, 4H), 3.20-2.80 (m, 5H), 2.60 (m, 1H), 2.20-1.60 (m, 5H). MS (EI) for C₃₆H₃₆N₈O₃: 629 (MH⁺).

[1403] 3-[5-Amino-6-(5-{2-[(phenylmethyl)oxy]ethyl}-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide: ¹H NMR (400 MHz, d₄-MeOH): δ 8.8 (s, 1H), 8.7 (s, 1H), 8.2 (m, 1H), 7.80 (m, 1H), 7.4 (m, 2H), 7.25 (m, 5H), 7.2 (m, 3H), 5.8 (m, 1H), 3.0 (m, 4H), 2.8 (m, 2H), 2.4 (m, 2H), 2.0 (m, 2H). MS (EI) for C₃₁H₂₉N₇O₂: 532 (MH⁺).

[1404] 3-[5-Amino-6-(1-phenyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide: ¹H NMR (400 MHz, d₄-MeOH): δ 8.72-8.61 (m, 2H), 8.22-8.17 (m, 1H), 7.97-7.91 (m, 1H), 7.70-7.56 (m, 6H), 7.36-7.15 (m, 4H), 5.76-5.66 (m, 1H), 3.58-3.46 (m, 2H), 3.15-2.86 (m, 4H), 2.68-2.57 (m, 1H), 2.15-1.75 (m, 6H). MS (EI) for C₃₃H₃₂N₈O: 557 (MH⁺).

[1405] 3-{5-Amino-6-[amino(imino)methyl]pyrazin-2-yl}-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide: ¹H NMR (400 MHz, d₆-DMSO): δ 8.95 (m, 2H), 8.50 (s, 1H), 8.20 (d, 1H), 7.95 (d, 1H), 7.60 (m, 1H), 7.40-7.20 (m, 4H), 5.60 (m, 1H), 3.20-2.80 (m, 2H), 2.20-2.00 (m, 2H). MS (EI) for C₂₁H₂₀N₆O: 373 (MH⁺).

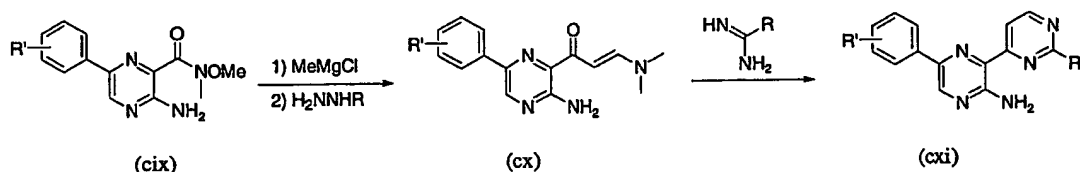
[1406] 3-[5-Amino-6-(5-methyl-1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide: ¹H NMR (400 MHz, d₆-DMSO): δ 9.00-8.90 (m, 3H), 8.50 (s, 1H), 8.20 (d, 1H), 7.95 (m, 2H), 7.60 (m, 2H), 7.40-7.20 (m, 4H), 5.60 (m, 1H), 3.20-2.80 (m, 2H), 2.75 (s, 3H), 2.20-2.00 (m, 2H). MS (EI) for C₂₈H₂₄N₈O: 489 (MH⁺).

- [1407] 3-[5-Amino-6-(1,5-dimethyl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]benzamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 8.75 (m, 2H), 8.20 (d, 1H), 7.95 (d, 1H), 7.60 (m, 1H), 7.40-7.20 (m, 4H), 5.70 (m, 1H), 4.00 (s, 3H), 3.20-2.80 (m, 2H), 2.60 (s, 3H), 2.20-2.00 (m, 2H). MS (EI) for C₂₄H₂₃N₇O: 426 (MH⁺).
- [1408] 3-[5-Amino-6-(1-pyridin-4-yl-1*H*-1,2,4-triazol-3-yl)pyrazin-2-yl]-*N*-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]benzamide: ¹H NMR (400 MHz, *d*₄-MeOH): δ 9.00 (s, 1H), 8.95 (m, 1H), 8.60 (s, 1H), 8.25 (d, 1H), 8.20-8.00 (m, 3H), 7.90-7.60 (m, 4H), 7.40-7.00 (m, 3H), 5.80 (m, 1H), 3.20-2.60 (m, 3H), 2.20-2.00 (m, 1H). MS (EI) for C₂₇H₂₂N₈O: 475 (MH⁺).

Example 71

- [1409] Scheme 27 depicts another strategy for making exemplary compounds according to formula I where A is a pyrimidine. Weinreb's amide intermediates (cix) were converted to α,β-unsaturated intermediates (cx), which were coupled with amidines and the intermediates cyclized to form pyrimidines (cxi). Intermediate (cix) can also be used to make other heterocycles for A, according to formula I.

Scheme 27



- [1410] 3-[5-Amino-6-(2-aminopyrimidin-4-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: To a THF (50 mL) solution of 3-amino-6-{3-[(benzylamino)carbonyl]phenyl}-*N*-methoxy-*N*-methylpyrazine-2-carboxamide (1.0 g, 2.6 mmol) was added 3.0 M methylmagnesium chloride in THF (6.1 mL, 18 mmol) and the mixture was stirred for 3 h. Saturated ammonium chloride was added and the mixture was extracted 3x with ethyl acetate. The combined organic layers were dried over sodium sulfate, filtered, and concentrated under reduced pressure. The crude product was purified by silica gel chromatography to afford the pure methyl ketone (0.400 g, 45% yield). This material was dissolved in ethanol (20 mL), dimethylformamide dimethyl acetal (2 mL, 15 mmol) was added and the mixture was heated to reflux for 90 min. LCMS analysis showed desired product and complete consumption of starting material. The reaction mixture was

concentrated under reduced pressure and redissolved in ethanol (10 mL). Guanidine (0.364 g, 2.0 mmol) was added followed by potassium carbonate (0.80 g, 5.8 mmol). The mixture was heated in a sealed tube at 80 °C overnight. The mixture was transferred to a round-bottomed flask and concentrated under reduced pressure. Water (100 mL) was added and the residue was extracted with ethyl acetate (3x 100 mL). The combined organic fractions were washed with water, dried over anhydrous sodium sulfate, filtered and concentrated under reduced pressure to give the desired product (0.103 g, 45% yield). ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.22 (br s, 1H), 8.86 (s, 1H), 8.56 (m, 1H), 8.40 (d, 1H), 8.25 (d, 1H), 7.90 (d, 1H), 7.71 (d, 1H), 7.59 (t, 1H), 7.52 (br s, 1H), 7.36 (m, 4H), 7.26 (m, 1H), 7.01 (br s, 1H), 4.53 (m, 3H). MS (EI) for C₂₂H₁₉N₇O: 398 (MH⁺).

[1411] Using the same or similar synthetic techniques, substituting with the appropriate reagents such as the respective amines, the following compounds of the invention were prepared:

[1412] 3-{5-Amino-6-[2-(dimethylamino)pyrimidin-4-yl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.20 (t, 1H), 8.80 (s, 1H), 8.57 (s, 1H), 8.54 (d, 1H), 8.26 (d, 1H), 7.90 (d, 1H), 7.75 (d, 1H), 7.59 (t, 1H), 7.36 (m, 4H), 7.26 (m, 1H), 4.58 (m, 1H), 4.54 (d, 2H), 3.20 (s, 6H). MS (EI) for C₂₄H₂₃N₇O: 426 (MH⁺).

[1413] 3-[5-Amino-6-(2-methylpyrimidin-4-yl)pyrazin-2-yl]-*N*-(phenylmethyl)benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.20 (t, 1H), 8.92 (s, 1H), 8.88 (d, 1H), 8.59 (s, 1H), 8.43 (d, 1H), 8.39 (m, 1H), 8.30 (d, 1H), 7.92 (d, 1H), 7.60 (t, 1H), 7.36 (m, 5H), 7.28 (m, 1H), 4.58 (t, 1H), 4.54 (d, 2H), 2.77 (s, 3H). MS (EI) for C₂₃H₂₀N₆O: 397 (MH⁺).

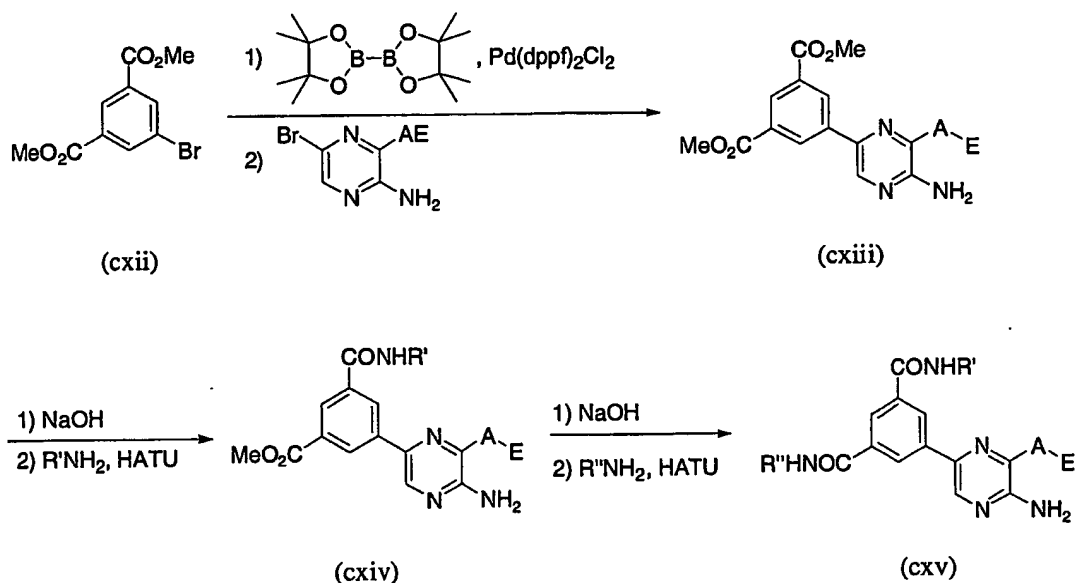
[1414] 3-{5-Amino-6-[2-(methylamino)pyrimidin-4-yl]pyrazin-2-yl}-*N*-(phenylmethyl)benzamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.20 (br s, 1H), 8.86 (m, 1H), 8.56 (m, 1H), 8.45 (m, 1H), 8.25 (m, 1H), 7.89 (m, 1H), 7.72 (m, 1H), 7.60 (m, 1H), 7.35 (m, 4H), 7.27 (m, 2H), 4.54 (m, 3H), 2.87 (m, 3H). MS (EI) for C₂₃H₂₁N₇O: 412 (MH⁺).

Example 72

[1415] Scheme 28 depicts another strategy for making exemplary compounds according to formula I where W is a substituted phenylene. Analogous to dimerization described in

Example 51, Scheme 17, above, intermediates (cxiii) were prepared by coupling aryl halides such as (cxii) to appropriate pyrazines, as depicted. In this example, mono-saponification of the esters and thus selective amide formation gave compounds (cxiv). The saponification process was repeated to make compounds (cxv).

Scheme 28



[1416] Dimethyl 5-{5-amino-6-[(methyloxy)carbonyl]pyrazin-2-yl}benzene-1,3-dicarboxylate: A mixture containing 5-bromoisophthalic acid dimethyl ester (0.20 g, 0.73 mmol), bispinacolatodiboron (0.21 g, 0.80 mmol), 1,1'-bis(diphenylphosphine)ferrocene palladium dichloride (0.029 g, 0.035 mmol), and potassium acetate (0.26 g, 2.3 mmol) in dioxane (5 mL) was heated at 80 °C for 1 h. The reaction mixture was cooled to room temperature and methyl 3-amino-6-bromo-pyrazine-2-carboxylate (0.17 mg, 0.70 mmol), 1,1'-bis(diphenylphosphine)ferrocene palladium dichloride (0.035 g, 0.042 mmol), sodium bicarbonate (320 mg, 3.8 mmol), and water (1 mL) were added. The mixture was heated at 80 °C for 12 h, after which time, the mixture was diluted with water and ethyl acetate. The layers were separated and the organic layer was extracted with 1 N hydrochloric acid (2x) and dried over sodium sulfate. Filtration, concentration, and silica gel chromatography gave the product as a yellow solid (23 mg, 12% yield). ^1H NMR (400 MHz, d_6 -DMSO): δ 9.04 (s, 1H), 8.74 (d, 2H), 8.44 (t, 1H), 7.62 (s, 2H), 3.32 (s, 3H), 2.49 (s, 3H), 2.48 (s, 3H). MS (EI) for $\text{C}_{16}\text{H}_{15}\text{N}_3\text{O}_6$: 346 (MH^+).

- [1417] 3-{5-Amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-5-[(methyloxy)carbonyl]benzoic acid: A mixture containing dimethyl 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzene-1,3-dicarboxylate (1.1 g, 3.2 mmol) and aqueous sodium hydroxide (3 mL of a 1N solution) was heated in dioxane (100 mL) at 95 °C for 3 h. The mixture was diluted with 1 N aqueous hydrochloric acid (300 mL) and extracted with ethyl acetate (3x 300 mL) and saturated ammonium chloride solution and dried over sodium sulfate. Filtration, concentration and purification by reverse phase HPLC gave the product as a yellow solid (0.39 g, 37% yield). ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.83 (s, 1H), 8.78 (d, 1H), 8.70 (d, 2H), 8.39 (t, 1H), 7.69 (br, 2H), 6.47 (br, 1H), 3.87 (s, 3H), 2.79 (d, 3H). MS (EI) for C₁₅H₁₄N₄O₅: 331 (MH⁺).
- [1418] Methyl 3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-5-[(1*S*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl}benzoate: A mixture containing 3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-5-[(methyloxy)carbonyl]benzoic acid (0.34 g, 1.0 mmol), O-(7-azabenzotriazol-1-yl)-N,N,N',N'-tetramethyluroniumhexafluorophosphate (0.51 g, 1.6 mmol), (S)-(+)-1-aminoindan (0.17 g, 1.3 mmol), diisopropylethylamine (0.6 mL, 3.4 mmol) and DMF (70 mL) were stirred at room temperature for 12 h. The reaction mixture was diluted with water and extracted 3x with ethyl acetate and 3x with 5% aqueous lithium chloride, and dried over sodium sulfate. Filtration, concentration, and purification by reverse phase HPLC gave the product as a yellow solid (0.41 g, 90% yield). ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.09 (d, 1H), 8.91 (s, 1H), 8.83 (d, 1H), 8.75 (t, 1H), 8.70 (t, 1H), 8.45 (t, 1H), 7.73 (br, 2H), 7.28 (d, 2H), 7.24-7.17 (m, 2H), 5.64-5.58 (q, 1H), 3.91 (s, 3H), 3.04-3.00 (m, 1H), 3.00-2.87 (m, 1H), 2.83 (d, 3H), 2.53-2.51 (m, 1H), 2.06-1.96 (m, 1H). MS (EI) for C₂₄H₂₃N₅O₄: 446 (MH⁺).
- [1419] 3-Amino-6-{3-[(2,3-dihydro-1*H*-inden-1-ylamino)carbonyl]-4-fluorophenyl}-*N*-methylpyrazine-2-carboxamide: A solution of methyl 2-fluoro-5-iodobenzoate (0.41 g, 1.5 mmol), bispinacolatodiboron (0.41 g, 1.6 mmol), Pd(dppfCl₂)·CH₂Cl₂ (180 mg, 0.22 mmol) and potassium acetate (0.43 g, 4.4 mmol) was heated at 80 °C in dioxane (50 mL) for 48 h. To this solution was added 3-amino-6-bromo-*N*-methylpyrazine-2-carboxamide (0.35 g, 1.5 mmol), additional Pd(dppfCl₂)·CH₂Cl₂ (0.2 g, 0.2 mmol), sodium bicarbonate (0.5 g, 6 mmol), and water (5mL) and the resulting solution was heated at 90 °C for an additional 16 h. The cooled solution was filtered through celite and concentrated under reduced pressure. Dichloromethane (100 mL) was added to the residue, which was washed with water, brine, and dried over anhydrous sodium sulfate. Filtration,

concentration and silica gel chromatography gave the biaryl product (0.24 g, 54% yield) as a light yellow solid. A solution of this material and lithium hydroxide (160 mg, 3.8 mmol) in dioxane (60 mL) and water (5 mL) was heated at 70 °C for 6 h. To the cooled solution was added 1M aqueous HCl and the reaction was concentrated to approximately 10 mL. The resulting solid was filtered and dried to yield the carboxylic acid (0.22 g, 97% yield) as a yellow solid. To a solution of this material, (S)-(+)-aminoindane (107 mg, 0.8 mmol) and Hunig's base (0.7 mL, 4 mmol) in DMF (10 mL) was added HATU (0.38 g, 1.0 mmol) and the solution was stirred at room temperature for 4 h. Ethyl acetate (50 mL) was added to the solution and it was washed with a 5% aqueous solution of lithium chloride (3x 50 mL) and brine, dried over anhydrous sodium sulfate, filtered, and concentrated under reduced pressure to yield a residue that was purified by silica gel chromatography to yield the desired product as a pale yellow solid (64mg, 20% yield). ¹H NMR (400 MHz, CDCl₃): δ 8.69-8.66 (m, 2H), 8.38 (m, 1H), 7.99-7.95 (m, 2H), 7.40-7.02 (m, 7H), 5.77-5.75 (m, 1H), 3.07-2.75 (m, 6H), 1.99-1.95 (m, 1H). MS (EI) for C₂₂H₂₀FN₅O₂: 406 (MH⁺).

[1420] Using the same or similar synthetic techniques, substituting with the appropriate reagents such as the respective amines, the following compounds of the invention were prepared:

[1421] 3-Amino-6-{3-[(2,3-dihydro-1*H*-inden-1-ylamino)carbonyl]-5-fluorophenyl}-*N*-methylpyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.96-8.93 (m, 3H), 8.39-8.29 (m, 2H), 7.66-7.64 (m, 1H), 7.29-7.19 (m, 4H), 5.59 (m, 1H), 3.34-2.84 (m, 6H), 2.03-1.98 (m, 1H). MS (EI) for C₂₂H₂₀FN₅O₂: 406 (MH⁺).

[1422] Dimethyl 5-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzene-1,3-dicarboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.85 (s, 1H), 8.79 (d, 1H), 8.73-8.72 (m, 2H), 8.39-8.38 (m, 1H), 7.75 (br, 1H), 3.87 (s, 6H), 3.82 (br, 1H), 2.79 (d, 3H). MS (EI) for C₁₆H₁₆N₄O₅: 345 (MH⁺).

[1423] 1,1-Dimethylethyl (3*S*)-3-[[[(3-amino-6-{3-[(2,3-dihydro-1*H*-inden-1-ylamino)sulfonyl]phenyl}pyrazin-2-yl)carbonyl]amino]pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.80 (s, 1H), 8.59 (d, 1H), 8.40-8.38 (m, 2H), 8.08 (br, 1H), 7.79 (t, 1H), 7.67 (br, 1H), 7.66-7.62 (m, 2H), 7.10-7.04 (m, 4H), 4.70 (t, 1H), 4.43-4.39 (m, 1H), 3.56-3.47 (m, 1H), 3.40-3.34 (m, 1H), 3.24-3.17 (m, 2H), 2.76-2.70 (m,

1H), 2.69-2.53 (m, 1H), 2.04-1.89 (m, 3H), 1.63-1.54 (m, 1H), 1.33 (s, 9H). MS (EI) for C₂₉H₃₄N₆O₅S: 577 (MH⁺).

[1424] 3-Amino-6-{3-[(2,3-dihydro-1*H*-inden-1-ylamino)sulfonyl]phenyl}-*N*-[(3*S*)-pyrrolidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.86 (s, 1H), 8.69-8.68 (m, 1H), 8.50-8.47 (m, 2H), 7.85 (d, 1H), 7.75 (br s, 1H), 7.70 (t, 2H), 7.14 (d, 4H), 4.77 (t, 1H), 4.41 (br s, 1H), 3.10-2.99 (m, 2H), 2.86-2.76 (m, 3H), 2.68-2.60 (m, 1H), 2.49 (s, 2H), 2.09-2.03 (m, 2H), 1.83-1.75 (m, 1H), 1.70-1.61 (m, 1H). MS (EI) for C₂₄H₂₆N₆O₃S: 477 (MH⁺).

[1425] 1,1-Dimethylethyl-3-{[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-5-{[(1*R*)-2,3-dihydro-1*H*-inden-1-ylamino]carbonyl}phenyl)carbonyl]amino}pyrrolidine-1-carboxylate: ¹H NMR (400 MHz, *d*₆-DMSO): δ 8.95 (d, 1H), 8.88 (s, 1H), 8.80 (d, 1H), 8.72 (d, 1H), 8.59 (s, 1H), 8.53 (s, 1H), 8.26 (s, 1H), 7.73 (br, 2H), 7.23 (d, 2H), 7.20-7.12 (m, 2H), 5.55 (q, 1H), 4.40 (m, 1H), 3.55-3.50 (m, 1H), 3.42-3.35 (m, 1H), 3.25-3.18 (m, 1H), 2.99-2.82 (m, 1H), 2.60 (m, 1H), 2.46-2.44 (m, 1H), 2.43 (s, 3H), 2.43-2.26 (m, 1H), 2.10-1.88 (m, 3H), 1.34 (s, 9H). MS (EI) for C₃₂H₃₇N₇O₅: 598 (MH⁺).

[1426] 5-{5-Amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-*N*-[(1*S*)-2,3-dihydro-1*H*-inden-1-yl]-*N'*-pyrrolidin-3-ylbenzene-1,3-dicarboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.02 (d, 1H), 8.93 (s, 1H), 8.84-8.80 (m, 4H), 8.67 (t, 1H), 8.60 (t, 1H), 8.33 (t, 1H), 7.77 (br s, 2H), 7.28 (d, 2H), 7.24-7.17 (m, 2H), 5.60 (q, 1H), 4.57-4.51 (m, 1H), 3.39 (m, 1H), 3.32-3.21 (m, 2H), 3.04-2.98 (m, 1H), 2.91-2.87 (m, 1H), 2.84 (d, 3H), 2.50 (m, 1H), 2.27-2.18 (m, 1H), 2.10-1.89 (s, 2H). MS (EI) for C₂₇H₂₉N₇O₃: 500 (MH⁺).

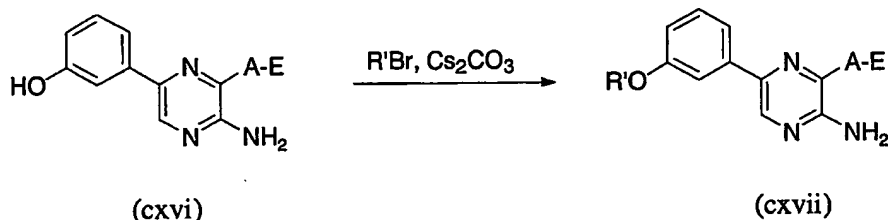
[1427] 5-{5-Amino-6-[(methylamino)carbonyl]pyrazin-2-yl}-*N*-(2,3-dihydro-1*H*-inden-1-yl)-*N'*-pyrrolidin-3-ylbenzene-1,3-dicarboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.02 (d, 1H), 8.93 (s, 1H), 8.85-8.81 (m, 4H), 8.67 (t, 1H), 8.61 (t, 1H), 8.34 (t, 1H), 7.28 (d, 2H), 7.24-7.19 (m, 2H), 7.56 (br, 2H), 5.60 (q, 1H), 4.56-4.50 (m, 1H), 3.50-3.35 (m, 2H), 3.32-3.18 (m, 2H), 3.04-3.97 (m, 1H), 2.91-2.87 (m, 1H), 2.84 (d, 3H), 2.27-2.18 (m, 1H), 2.10-2.96 (m, 2H). MS (EI) for C₂₇H₂₉N₇O₃: 500 (MH⁺).

Example 73

[1428] Scheme 29 depicts another strategy for making exemplary compounds according to formula I where W is a substituted phenylene, specifically where W possessed a phenolic

group and it was desirable to selectively alkylate the phenolic oxygen. Compounds (cxvii) were prepared by coupling phenols with, for example, alkyl bromides, as depicted.

Scheme 29



[1429] 3-Amino-6-(3-{[2-(4-phenylpiperazin-1-yl)ethyl]oxy}phenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: A mixture of 3-amino-6-(3-hydroxyphenyl)-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide (0.11 g, 0.27 mmol), 1,2-dibromoethane (4 mL, 46 mmol), and cesium carbonate (0.35 g, 1.1 mmol) in DMF (2 mL) was heated at 80 °C for 12 h. The reaction mixture was cooled to room temperature, diluted with 1 N aqueous hydrochloric acid solution, and extracted 3x with ethyl acetate, 2x with 5% aqueous lithium chloride, and dried over sodium sulfate. Filtration, concentration, and silica gel chromatography gave the corresponding ether. A mixture of this ether intermediate (0.012 g, 0.023 mmol) and *N*-phenylpiperazine (4.4 mg, 0.027 mmol) was stirred in acetonitrile (5 mL) at room temperature for 3 h. Concentration followed by reverse phase HPLC gave the protected product as a yellow solid. This solid was added to 4 N hydrochloric acid in dioxane (5 mL) and the mixture was stirred at room temperature for 1 h. Removal of solvent under reduced pressure gave the desired product as a yellow solid (3.2 mg, 28% yield). ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.10 (br s, 1H), 8.88 (s, 1H), 8.71 (br, 1H), 7.78-7.67 (m, 2H), 7.65 (br s, 1H), 7.42 (t, 1H), 7.25 (t, 2H), 7.04-6.98 (m, 3H), 6.85 (t, 1H), 4.56 (br, 2H), 4.24 (br, 2H), 3.83 (d, 2H), 3.70-3.63 (m, 4H), 3.48-3.45 (m, 1H), 3.30-3.18 (m, 6H), 2.76 (br, 1H), 1.87-1.78 (m, 4H). MS (EI) for C₂₈H₃₅N₇O₂: 502 (MH⁺).

[1430] Using the same or similar synthetic techniques, substituting with the appropriate reagents such as the respective amines, the following compounds of the invention were prepared:

[1431] 3-Amino-6-(3-{[2-(2-hydroxyethyl)oxy]phenyl}-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide: ¹H NMR (400 MHz, *d*₆-DMSO): δ 9.15-9.08 (m, 1H), 8.95-8.92 (m, 1H), 8.85 (s, 1H), 8.69 (d, 1H), 7.66 (br s, 1H), 7.69 (d, 1H), 7.63 (s, 1H), 7.36 (t, 1H), 6.95 (dd, 1H), 4.22 (br s, 1H), 4.07 (t, 2H), 3.23 (t, 2H), 3.05-3.02 (m, 1H), 2.76-2.74 (m, 1H),

1.97-1.88 (m, 2H), 1.76-1.74 (m, 2H), 1.26-1.13 (m, 2H), 0.87-0.81 (m, 1H). MS (EI) for $C_{18}H_{23}N_5O_3$: 358 (MH^+).

[1432] 3-Amino-6-[3-(methyloxy)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-

carboxamide: 1H NMR (400 MHz, d_6 -DMSO): δ 9.02-8.99 (m, 1H), 8.86 (s, 1H), 8.85 (m, 1H), 8.68 (d, 1H), 7.70-7.65 (m, 3H), 7.37 (t, 1H), 6.95 (dd, 1H), 4.23 (m, 1H), 3.82 (s, 3H), 3.27-3.19 (m, 2H), 3.07-2.99 (m, 1H), 2.77-2.74 (m, 1H), 1.89-1.87 (m, 2H), 1.76-1.72 (m, 2H). MS (EI) for $C_{17}H_{21}N_5O_2$: 328 (MH^+).

[1433] 3-Amino-6-[3-(ethyloxy)phenyl]-*N*-[(3*S*)-piperidin-3-yl]pyrazine-2-carboxamide:

1H NMR (400 MHz, d_6 -DMSO): δ 9.03-9.00 (m, 1H), 8.87-8.82 (m, 1H), 8.81 (s, 1H), 8.64 (d, 1H), 7.64-7.57 (m, 3H), 7.31 (t, 1H), 6.89 (dd, 1H), 4.19-4.17 (m, 1H), 4.06 (q, 2H), 3.22-3.15 (m, 2H), 3.02-2.95 (m, 1H), 2.72-2.70 (m, 1H), 1.85-1.83 (m, 2H), 1.71-1.68 (m, 2H), 1.30 (t, 3H). MS (EI) for $C_{18}H_{23}N_5O_2$: 342 (MH^+).

[1434] 3-Amino-6-(3-{[2-(diethylamino)ethyl]oxy}phenyl)-*N*-[(3*S*)-piperidin-3-

yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.63 (s, 1H), 7.64-7.62 (m, 2H), 7.36 (t, 1H), 6.98 (d, 1H), 4.40 (br s, 2H), 4.25 (br s, 1H), 3.66-3.63 (m, 2H), 3.59-3.51 (m, 9H), 3.50-3.47 (m, 2H), 1.95 (br s, 2H), 1.83 (br, 2H), 1.31 (m, 7H). MS (EI) for $C_{22}H_{32}N_6O_2$: 413 (MH^+).

[1435] 3-Amino-6-(3-{[2-(morpholin-4-ylethyl)oxy]phenyl})-*N*-[(3*S*)-piperidin-3-

yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (br s, 1H), 7.80 (br s, 1H), 7.71 (d, 1H), 7.45 (m, 1H), 7.09 (d, 1H), 4.55 (m, 2H), 4.36 (m, 1H), 4.10-4.06 (m, 2H), 3.93 (m, 2H), 3.74-3.71 (m, 2H), 3.67-3.64 (m, 9H), 3.58-3.56 (m, 2H), 3.48-3.47 (m, 1H), 2.10-2.04 (m, 2H), 1.89 (m, 2H). MS (EI) for $C_{22}H_{30}N_6O_3$: 427 (MH^+).

[1436] 3-Amino-6-(3-{[2-(ethylamino)ethyl]oxy}phenyl)-*N*-[(3*S*)-piperidin-3-

yl]pyrazine-2-carboxamide: 1H NMR (400 MHz, d_4 -MeOH): δ 8.70 (s, 1H), 7.73-7.70 (m, 2H), 7.44 (t, 1H), 7.08 (d, 1H), 4.41-4.34 (m, 2H), 3.75-3.72 (m, 2H), 3.70-3.63 (m, 5H), 3.58-3.56 (m, 1H), 3.50 (br s, 2H), 3.37-3.34 (m, 1H), 3.21-3.19 (m, 2H), 3.04 (m, 1H), 2.10-2.04 (m, 2H), 1.98-1.92 (m, 2H), 1.38 (t, 3H). MS (EI) for $C_{20}H_{28}N_6O_2$: 385 (MH^+).

[1437] Phenylmethyl 4-(2-{[3-(5-amino-6-{[(3*S*)-piperidin-3-ylamino]carbonyl}pyrazin-

2-yl)phenyl]oxy}ethyl)piperazine-1-carboxylate: 1H NMR (400 MHz, d_4 -MeOH): δ 8.69 (br s, 1H), 7.87 (br s, 1H), 7.70 (m, 1H), 7.45 (m, 1H), 7.37-7.31 (m, 4H), 7.10 (d, 1H), 5.16 (s, 2H), 4.56 (m, 2H), 4.37-4.28 (m, 4H), 3.74-3.70 (m, 4H), 3.69-3.62 (m, 4H),

3.58-3.56 (m, 2H), 3.34 (m, 4H), 3.05 (m, 2H), 2.08 (m, 2H), 1.89 (m, 2H). MS (EI) for $C_{30}H_{37}N_7O_4$: 560 (MH^+).

[1438] 3-Amino-N-[(3S)-azepan-3-yl]-6-{3-[[[4-(1H-pyrazol-1-yl)phenyl]methyl]amino)carbonyl]phenyl}pyrazine-2-carboxamide: 1H NMR (400 MHz, DMSO- d_6): δ 9.59 (m, 2H), 9.25 (bs, 1H), 9.08 (d, 1H), 8.96 (s, 1H), 8.78 (m, 1H), 8.48 (d, 1H), 8.32 (m, 1H), 7.92 (m, 1H), 7.80 (m, 1H), 7.78 (m, 1H), 7.73 (m, 1H), 7.58 (t, 1H), 7.50 (d, 2H), 6.53 (m, 1H), 4.77 (bs, 2H), 4.53 (m, 2H), 4.43 (m, 1H), 3.34 (m, 2H), 3.24 (m, 1H), 2.99 (m, 1H), 2.01 (m, 2H), 1.80 (m, 3H), 1.54 (m, 1H); MS (EI) for $C_{28}H_{30}N_8O_2$ 511.36 (MH^+).

Assays

[1439] Chk1 and Chk2 assays were performed by measuring incorporation of γ - ^{33}P ATP into immobilized myelin basic protein (MBP). High binding white 384 well plates (Greiner) were coated with MBP (Sigma #M-1891) by incubation of 60 μ l/well of 20 μ g/ml MBP in Tris-buffered saline (TBS; 50mM Tris pH 8.0, 138mM NaCl, 2.7mM KCl) for 24 hours at 4° C. Plates were washed 3X with 100 μ l TBS. Kinase reactions were carried out in a total volume of 34 μ l in kinase buffer (5mM Hepes pH 7.6, 15mM NaCl, 0.01% bovine gamma globulin (Sigma #I-5506), 10mM $MgCl_2$, 1mM DTT, 0.02% TritonX-100). Compound dilutions were performed in DMSO and added to assay wells to a final DMSO concentration of 1%. Each data point was measured in duplicate, and at least two duplicate assays were performed for each individual compound determination. Chk1 and Chk2 enzymes (Upstate Biotechnology) were added to final concentrations of 10nM (Chk1) or 20nM (Chk2). A mixture of unlabeled ATP and γ - ^{33}P ATP was added to start the reaction (2×10^6 cpm of γ - ^{33}P ATP per well (3000Ci/mmol) and either 10 μ M or 30 μ M unlabeled ATP (for Chk1 and Chk2 assays respectively). The reaction was carried out for 1 hour at room temperature with shaking. Plates were washed 7x with TBS, followed by the addition of 50 μ l/well scintillation fluid (Wallac). Plates were read using a Wallac Trilux counter. This is only one format of such assays, various other formats are possible, as known to those skilled in the art.

[1440] The above assay procedure can be used to determine the IC_{50} for inhibition and/or the inhibition constant K_i . The IC_{50} is defined as the concentration of compound required to reduce the enzyme activity by 50% under the conditions of the assay. Exemplary

compositions have IC₅₀'s of, for example, less than about 100 μM, less than about 10 μM, less than about 1 μM, and further for example having IC₅₀'s of less than about 100 nM, and still further, for example, less than about 10 nM. The K_i for a compound may be determined from the IC₅₀ based on three assumptions. First, only one compound molecule binds to the enzyme and there is no cooperativity. Second, the concentrations of active enzyme and the compound tested are known (i.e., there are no significant amounts of impurities or inactive forms in the preparations). Third, the enzymatic rate of the enzyme-inhibitor complex is zero. The rate (i.e., compound concentration) data are fitted to the equation:

$$V = V_{\max} E_0 \left[I - \frac{(E_0 + I_0 + K_d) - \sqrt{(E_0 + I_0 + K_d)^2 - 4E_0 I_0}}{2E_0} \right]$$

[1441] Where V is the observed rate, V_{max}, is the rate of the free enzyme, I₀ is the inhibitor concentration, E₀ is the enzyme concentration, and K_d is the dissociation constant of the enzyme-inhibitor complex.

Kinase specificity assays:

[1442] Kinase activity and compound inhibition are investigated using one or more of the three assay formats described below. The ATP concentrations for each assay are selected to be close to the Michaelis-Menten constant (K_M) for each individual kinase. Dose-response experiments are performed at 10 different inhibitor concentrations in a 384-well plate format. The data are fitted to the following four-parameter equation:

$$Y = \text{Min} + (\text{Max} - \text{Min}) / (1 + (X/\text{IC}_{50})^H)$$

where Y is the observed signal, X is the inhibitor concentration, Min is the background signal in the absence of enzyme (0% enzyme activity), Max is the signal in the absence of inhibitor (100% enzyme activity), IC₅₀ is the inhibitor concentration at 50% enzyme inhibition and H represents the empirical Hill's slope to measure the cooperativity. Typically H is close to unity.

³³P phosphoryl transfer assay:

[1443] Greiner 384-well white, clear bottom, high binding plates are coated with 2 µg/well of protein or peptide substrate in a 50 µL volume overnight at ambient temperature. The coating buffer typically contains 40 µg/mL substrate, 22.5 mM Na₂CO₃, 27.5 mM NaHCO₃, 150 mM NaCl and 3 mM NaN₃. The coating solution is aspirated and the plates are washed once with 50 µL of assay buffer and padded dry. Subsequently compounds and enzymes are mixed with γ ³³P-ATP (3.3µCi/nmol) in a total volume of 20 uL in suitable assay buffers. The mixture is incubated at ambient temperature for 1.5-2.5 hrs and terminated by aspiration using a Molecular Devices EMBLA 96-head plate washer. The plates are subsequently washed 6-12 times with PBST or TBS buffer. Scintillation fluid (50µl/well) is then added, the plates are sealed and activity assessed by liquid scintillation spectrometry using a Perkin Elmer MicroBeta TriLux.

Luciferase-coupled chemiluminescent assay (LCCA)

[1444] In the LCCA assays, kinase activity is measured as the percent ATP consumed following the kinase reaction using luciferase-luciferin-coupled chemiluminescence. Greiner 384-well white, clear bottom, medium binding plates are used for LCCA. Briefly, the kinase reaction is initiated by mixing compounds, ATP and kinase in a 20 uL volume. The mixture is incubated at ambient temperature for 2-4 hrs. At the end of the kinase reaction, a 20 uL luciferase-luciferin mix is added and the chemiluminescent signal is read using a Wallac Victor² reader. The luciferase-luciferin mix consists of 50 mM HEPES, pH 7.8, 8.5 ug/mL oxalic acid (pH 7.8), 5 (or 50) mM DTT, 0.4% Triton X-100, 0.25 mg/mL coenzyme A, 63 uM AMP, 28 ug/mL luciferin and 40,000 units of light/mL luciferase. For the LCCA assays, total ATP consumption has been standardized to 25-45% and the IC₅₀ values correlates well with those determined by radiometric assays.

AlphaScreen

[1445] AlphaScreen technology is a proximity assay which utilizes the increase in fluorescent signal when donor and acceptor beads are close in proximity. Biotinylated poly(Glu,Tyr) 4:1 is used as the kinase substrate, in conjunction with streptavidin-coated donor beads and anti-phosphotyrosine antibody PY100-coated acceptor beads. Upon phosphorylation, the peptide substrate can bind to both donor and acceptor beads giving

rise to an increase in fluorescence. Compounds, ATP, biotinylated poly(Glu, Tyr) and kinases are mixed in a volume of 20 uL for 1 hr at ambient temperature using Greiner 384-well white clear bottom medium binding plates. Following incubation a 10 uL solution containing 15-30 mg/mL AlphaScreen beads, 75 mM Hepes, pH 7.4, 300 mM NaCl, 120 mM EDTA, 0.3% BSA and 0.03% Tween-20 is added to each well. After 2-16 hr incubation of the beads, plates are read in a Perkin Elmer AlphaQuest reader. The IC₅₀ values correlates well with those of radiometric assays.

[1446] In order to assess the ability of the compound to abrogate the G2 checkpoint in cells (*via* Chk1 inhibition), a mitotic index assay can be performed. Mitotic index was determined by measuring accumulation of cells in mitosis in the presence of a DNA damaging agent (for example, Adriamycin; ADR). Nocodazole (Noc) was used to induce mitotic arrest. In the presence of Noc, cells accumulate in mitosis and are detected using an M-phase specific antibody. In the presence of ADR and Noc, cells undergo G2 arrest and do not proceed to M-phase. Addition of Chk1 inhibitors causes the G2 checkpoint to be bypassed, and M-phase cells are detected. HT-29 cells were cultured by 1:20 passaging in DMEM + 10% fetal bovine serum (DMEM/FBS) at 37° C and 5% CO₂. Prior to the assay, cells were plated at 6500 cells/well in 100µl DMEM/FBS in black/clear bottom 96 well plates (BD Biocoat). After growing for 24 hours at 37° C/5% CO₂, ADR was added to a final concentration of 250nM and cells were grown for a further 18-20 hrs. Medium was then aspirated from the plates and 70µl of 0.71µM Noc added in DMEM/FBS containing 0.55% v/v DMSO. Compounds were diluted in DMEM/FBS containing 0.55% v/v DMSO, and 30µl of compound solution added to the assay plates. Control wells with Noc, compound/Noc and Noc/ADR alone were also set up. Cells were grown for a further 18-20 hrs and visually inspected before fixing with formaldehyde. The percentage of mitotic cells was determined by immunofluorescence using a M-phase specific anti-phosphohistoneH3 rabbit polyclonal antibody and a goat anti-rabbit secondary antibody conjugated to Alexa Fluor 488 dye. Cell nuclei were also stained using Hoechst dye. Staining and fluorescence detection was performed using a Cellomics Arrayscan instrument, using the manufacturer's mitotic index assay kit. This is only one format of such assays, various other formats are possible, as known to those skilled in the art.

Structure Activity Relationships

[1447] Table 11 shows structure activity relationship data for selected compounds of the invention with respect to Chk1 only. Inhibition is indicated as IC₅₀ with following key: A = IC₅₀ less than 50 nM, B = IC₅₀ greater than 50 nM but less than or equal to 1000 nM, C = IC₅₀ greater than 1000 nM but less than 10,000 nM, and D = IC₅₀ of 10,000 nM or greater.

Table 11

#	Name	Chk1 IC ₅₀
1	3-amino-6-phenyl-N-(2-phenylethyl)pyrazine-2-carboxamide	D
2	3-amino-6-phenyl-N-(phenylmethyl)pyrazine-2-carboxamide	D
3	3-amino-6-(3-chlorophenyl)pyrazine-2-carboxamide	D
4	3-amino-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide	D
5	3-amino-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide	D
6	3-amino-N-(2,3-dihydro-1H-inden-2-yl)-6-phenylpyrazine-2-carboxamide	D
7	3-amino-6-(2-chlorophenyl)pyrazine-2-carboxamide	D
8	3-amino-N-cyclopentyl-6-phenylpyrazine-2-carboxamide	D
9	3-amino-N,N-dimethyl-6-phenylpyrazine-2-carboxamide	C
10	3-amino-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide	C
11	5-phenyl-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine	D
12	3-amino-N-(cyclopropylmethyl)-6-phenylpyrazine-2-carboxamide	D
13	3-amino-6-phenyl-N-(tetrahydrofuran-2-ylmethyl)pyrazine-2-carboxamide	D
14	3-amino-N-cyclohexyl-6-phenylpyrazine-2-carboxamide	D
15	6-phenyl-N-propyl-3-(propylamino)pyrazine-2-carboxamide	D
16	3-amino-6-(2-methylphenyl)pyrazine-2-carboxamide	D
17	3-amino-6-[2-(methyloxy)phenyl]pyrazine-2-carboxamide	D
18	3-amino-6-(2,4-difluorophenyl)pyrazine-2-carboxamide	D

#	Name	Chk1 IC ₅₀
19	3-amino-6-(3-fluorophenyl)pyrazine-2-carboxamide	D
20	3-amino-N-cyclopropyl-6-phenylpyrazine-2-carboxamide	D
21	3-amino-6-(3,4-difluorophenyl)pyrazine-2-carboxamide	D
22	3-amino-6-(4-fluorophenyl)-N-methylpyrazine-2-carboxamide	C
23	3-amino-6-[4-(ethyloxy)phenyl]pyrazine-2-carboxamide	C
24	3-amino-6-[3-(ethyloxy)phenyl]pyrazine-2-carboxamide	C
25	3-amino-6-(1,3-benzodioxol-5-yl)pyrazine-2-carboxamide	D
26	3-amino-6-naphthalen-1-ylpyrazine-2-carboxamide	D
27	3-amino-6-naphthalen-2-ylpyrazine-2-carboxamide	D
28	3-amino-6-[3-[(trifluoromethyl)oxy]phenyl]pyrazine-2-carboxamide	C
29	3-amino-6-[3-(aminocarbonyl)phenyl]pyrazine-2-carboxamide	C
30	6-[3-(acetyl amino)phenyl]-3-aminopyrazine-2-carboxamide	C
31	3-amino-6-biphenyl-4-ylpyrazine-2-carboxamide	D
32	3-amino-6-[4-(dimethylamino)phenyl]pyrazine-2-carboxamide	C
33	3-amino-6-(3-methylphenyl)pyrazine-2-carboxamide	C
34	methyl 3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoate	C
35	3-amino-6-[3-[(phenylmethyl)oxy]phenyl]pyrazine-2-carboxamide	B
36	3-amino-6-(3-hydroxyphenyl)pyrazine-2-carboxamide	B
37	3-amino-6-(1-benzofuran-2-yl)pyrazine-2-carboxamide	D
38	3-(methylamino)-6-phenylpyrazine-2-carboxamide	D
39	6-phenyl-3-[(phenylmethyl)amino]pyrazine-2-carboxamide	D
40	6-phenyl-3-(propylamino)pyrazine-2-carboxamide	D
41	3-amino-6-biphenyl-3-ylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
42	3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoic acid	D
43	3-amino-6-{4-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide	D
44	3-amino-N-methyl-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide	C
45	3-amino-6-(3-hydroxyphenyl)-N-methylpyrazine-2-carboxamide	B
46	3-amino-N-[2-(methyloxy)ethyl]-6-phenylpyrazine-2-carboxamide	C
47	N-[2-(acetilamino)ethyl]-3-amino-6-phenylpyrazine-2-carboxamide	D
48	3-amino-6-phenylpyrazine-2-carbohydrazide	C
49	3-amino-N-hydroxy-6-phenylpyrazine-2-carboxamide	C
50	3-amino-6-[3-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide	C
51	3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide	B
52	3-amino-6-[4-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide	C
53	3-amino-N-methyl-6-(3-methylphenyl)pyrazine-2-carboxamide	B
54	3-amino-N-methyl-6-[4-(methyloxy)phenyl]pyrazine-2-carboxamide	C
55	3-amino-6-biphenyl-3-yl-N-methylpyrazine-2-carboxamide	B
56	6-[3-(acetilamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide	B
57	3-amino-6-[3-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide	B
58	3-amino-6-[4-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide	C
59	3-amino-N-methyl-6-{3-[(1E)-N-methylethanimidoyl]phenyl}pyrazine-2-carboxamide	C
60	3-amino-N-methyl-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide	C
61	3-amino-6-(4-fluoro-3-methylphenyl)-N-methylpyrazine-2-carboxamide	B
62	3-amino-N-methyl-6-pyridin-3-ylpyrazine-2-carboxamide	C
63	6-[4-(acetilamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide	C
64	3-amino-N-methyl-6-{3-[(methylamino)carbonyl]phenyl}pyrazine-2-carboxamide	C

#	Name	Chk1 IC ₅₀
65	6-(3-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide	C
66	3-amino-N-methyl-6-(5-pyridin-3-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide	D
67	3-amino-6-(5-cyclopentyl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide	D
68	3-amino-6-(5-cyclohexyl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide	D
69	3-amino-N-methyl-6-(5-phenyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide	D
70	3-amino-N-methyl-6-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide	D
71	3-amino-N-methyl-6-(5-methyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide	C
72	3-amino-N-methyl-6-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide	D
73	3-amino-N-methyl-6-(5-pyridin-4-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide	D
74	3-amino-6-[5-(1,1-dimethylethyl)-1,2,4-oxadiazol-3-yl]-N-methylpyrazine-2-carboxamide	D
75	3-amino-6-(5-furan-2-yl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide	D
76	3-amino-6-[4-(dimethylamino)phenyl]-N-methylpyrazine-2-carboxamide	C
77	3-amino-N-methyl-6-[4-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide	C
78	3-amino-6-(3,5-dimethylphenyl)-N-methylpyrazine-2-carboxamide	C
79	6-(4-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide	C
80	3-amino-6-[3,4-bis(methyloxy)phenyl]-N-methylpyrazine-2-carboxamide	C
81	3-amino-N-methyl-6-[3-[(phenylcarbonyl)amino]phenyl]pyrazine-2-carboxamide	B
82	3-amino-N-methyl-6-[3-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide	C
83	3-amino-N-methyl-6-[3-(1H-tetrazol-5-yl)phenyl]pyrazine-2-carboxamide	C
84	3-amino-N-methyl-6-[3-[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
85	3-amino-6-[3-(aminocarbonyl)phenyl]-N-methylpyrazine-2-carboxamide	C
86	3-amino-6-[3-[(dimethylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	C
87	3-amino-N-methyl-6-[4-(methylsulfonyl)phenyl]pyrazine-2-carboxamide	C

#	Name	Chk1 IC ₅₀
88	3-amino-6-(3-aminophenyl)-N-ethylpyrazine-2-carboxamide	C
89	3-amino-6-[3-(((4-fluorophenyl)amino)carbonyl)amino]phenyl]-N-methylpyrazine-2-carboxamide	B
90	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)amino)carbonyl)amino]phenyl]pyrazine-2-carboxamide	B
91	6,6'-[(oxomethanediy)bis(iminobenzene-3,1-diyl)]bis(3-amino-N-methylpyrazine-2-carboxamide)	A
92	3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoic acid	D
93	3-amino-N-methyl-6-[3-[(4-methylpiperazin-1-yl)carbonyl]phenyl]pyrazine-2-carboxamide	C
94	3-amino-N-methyl-6-[3-[(2-phenylethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
95	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
96	3-amino-6-[3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	C
97	3-amino-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
98	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
99	3-amino-N-methyl-6-[3-({[methyl(phenylmethyl)amino]carbonyl}phenyl)pyrazine-2-carboxamide	C
100	3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
101	3-amino-N-methyl-6-[3-(((4-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
102	3-amino-N-methyl-6-[3-(((4-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
103	3-amino-6-[3-(((4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
104	3-amino-6-[3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
105	3-amino-N-methyl-6-[3-(((3-(methyloxy)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide	B
106	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide	B
107	3-amino-N-methyl-6-[3-(((3-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
108	3-amino-N-methyl-6-[3-(((1R)-1-[4-(methyloxy)phenyl]ethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
109	3-amino-N-methyl-6-[3-(((2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
110	3-amino-6-[3-(((3-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
111	3-amino-6-(3-[[[4-chlorophenyl]carbonyl]amino]phenyl)-N-methylpyrazine-2-carboxamide	B
112	3-amino-6-[3-(((1-(4-fluorophenyl)ethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	C
113	3-amino-6-[3-(((2,3-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
114	3-amino-6-(3-[[bis(phenylmethyl)amino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide	C
115	3-amino-6-[3-(((2,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
116	3-amino-6-(3-[[ethyl(pyridin-4-ylmethyl)amino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide	C
117	3-amino-N-methyl-6-(3-[[4-(phenylmethyl)piperazin-1-yl]carbonyl]phenyl)pyrazine-2-carboxamide	C
118	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)piperidin-4-yl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
119	3-amino-N-methyl-6-(3-[[[4-methylphenyl]carbonyl]amino]phenyl)pyrazine-2-carboxamide	B
120	3-amino-N-methyl-6-[3-(((5-methylpyrazin-2-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
121	3-amino-N-methyl-6-(3-[[pyridin-2-ylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide	B
122	3-amino-N-methyl-6-(3-[[pyridin-3-ylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide	B
123	3-amino-N-methyl-6-(3-[[pyridin-4-ylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide	B
124	3-amino-6-[3-[[furan-2-ylcarbonyl]amino]phenyl]-N-methylpyrazine-2-carboxamide	B
125	3-amino-6-[3-(((3-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
126	3-amino-6-(3-[[cyclohexylmethyl]amino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide	B
127	3-amino-N-methyl-6-(3-[[2-phenylpropyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide	B
128	3-amino-6-[3-(((2-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
129	3-amino-6-[3-(((2-(2-chlorophenyl)ethyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide	B
130	3-amino-N-methyl-6-[3-(((2-(2-(methyloxy)phenyl)ethyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide	B
131	3-amino-6-[3-(((2-(4-fluorophenyl)ethyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide	B
132	3-amino-N-methyl-6-[3-(((2-(2-thienyl)ethyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide	B
133	3-amino-6-[3-[(cyclohexylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	C

#	Name	Chk1 IC ₅₀
134	1,1-dimethylethyl 4-(((3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbonyl)amino)methyl)piperidine-1-carboxylate	B
135	3-amino-N-methyl-6-[3-(((piperidin-4-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
136	3-amino-N-methyl-6-[3-(5-phenyl-1,2,4-oxadiazol-3-yl)phenyl]pyrazine-2-carboxamide	C
137	3-amino-N-methyl-6-[3-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]phenyl]pyrazine-2-carboxamide	C
138	3-amino-N-methyl-6-[3-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]phenyl]pyrazine-2-carboxamide	C
139	3-amino-6-[3-(((1,1-dimethylethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	C
140	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
141	3-amino-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	C
142	3-amino-N-methyl-6-[3-[(phenylamino)carbonyl]phenyl]pyrazine-2-carboxamide	C
143	3-amino-6-[3-(1,3-dihydro-2H-isindol-2-ylcarbonyl)phenyl]-N-methylpyrazine-2-carboxamide	C
144	3-amino-N-methyl-6-[3-(((1R)-1-phenylethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
145	3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
146	3-amino-N-methyl-6-[3-(((1R)-1-phenylpropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
147	3-amino-N-methyl-6-[3-(((1S)-1-phenylpropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
148	3-amino-N-methyl-6-[3-(((1R,2S)-2-phenylcyclopropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
149	phenylmethyl 4-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}benzoate	D
150	3-amino-N-methyl-6-[4-(((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	C
151	3-amino-N-methyl-6-[4-(((2-phenylethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
152	3-amino-6-[4-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
153	3-amino-N-methyl-6-[4-(((4-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
154	3-amino-6-[4-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
155	3-amino-6-[3-(((4-chlorophenyl)acetyl)amino)phenyl]-N-methylpyrazine-2-carboxamide	B
156	3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
157	3-amino-6-{3-[(diphenylacetyl)amino]phenyl}-N-methylpyrazine-2-carboxamide	B
158	3-amino-N-methyl-6-{3-[(phenylacetyl)amino]phenyl}pyrazine-2-carboxamide	B
159	3-amino-N-methyl-6-{3-[(3-phenylpropanoyl)amino]phenyl}pyrazine-2-carboxamide	B
160	3-amino-N-methyl-6-{3-[(1R)-1,2,3,4-tetrahydronaphthalen-1-ylamino]carbonyl}phenyl}pyrazine-2-carboxamide	C
161	3-amino-N-methyl-6-{3-[(1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino]carbonyl}phenyl}pyrazine-2-carboxamide	B
162	3-amino-N-methyl-6-{3-[[[3,4,5-tris(methyloxy)phenyl]methyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide	B
163	3-amino-N-methyl-6-{3-[[[(1R,2R)-2-[(phenylmethyl)oxy]cyclopentyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide	C
164	3-amino-N-methyl-6-{3-[[[(1S,2S)-2-[(phenylmethyl)oxy]cyclopentyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide	B
165	3-amino-6-(3-cyanophenyl)-N-methylpyrazine-2-carboxamide	C
166	3-amino-6-{3-[(cyclopentylcarbonyl)amino]phenyl}-N-methylpyrazine-2-carboxamide	B
167	3-amino-6-{3-[[4-cyanophenyl]carbonyl]amino]phenyl}-N-methylpyrazine-2-carboxamide	B
168	3-amino-N-methyl-6-{3-[[[(phenylmethyl)amino]carbonyl]amino]phenyl}pyrazine-2-carboxamide	B
169	3-amino-N-methyl-6-{3-[(naphthalen-2-ylmethyl)amino]carbonyl}phenyl}pyrazine-2-carboxamide	B
170	3-amino-N-methyl-6-{3-[[tetrahydrofuran-2-ylmethyl]amino]carbonyl}phenyl}pyrazine-2-carboxamide	C
171	3-amino-6-{3-[[2-chloro-6-fluorophenyl]methyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide	B
172	3-amino-6-{3-[[[3-fluoro-5-(trifluoromethyl)phenyl]methyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide	B
173	3-amino-6-{3-[[2-{3,5-bis(methyloxy)phenyl}ethyl]amino]carbonyl}phenyl}-N-methylpyrazine-2-carboxamide	C
174	3-amino-N-methyl-6-{3-[(2-phenylethyl)oxy]phenyl}pyrazine-2-carboxamide	B
175	3-amino-N-methyl-6-{3-[[2-(4-methylpiperidin-1-yl)ethyl]oxy]phenyl}pyrazine-2-carboxamide	C
176	3-amino-6-{3-[[2-{4-(2-hydroxyethyl)piperidin-1-yl}ethyl]oxy]phenyl}-N-methylpyrazine-2-carboxamide	C
177	3-amino-6-{3-[[2-(3-hydroxypyrrolidin-1-yl)ethyl]oxy]phenyl}-N-methylpyrazine-2-carboxamide	C
178	3-amino-6-{3-[(2-[ethyl(phenylmethyl)amino]ethyl)oxy]phenyl}-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
179	3-amino-6-(3-([2-(diethylamino)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide	B
180	3-amino-6-(3-([(biphenyl-2-ylmethyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	A
181	3-amino-N-methyl-6-(3-([2-(4-phenylpiperazin-1-yl)ethyl]oxy)phenyl)pyrazine-2-carboxamide	B
182	3-amino-6-(3-([(biphenyl-4-ylmethyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	A
183	3-amino-N-methyl-6-[3-([2-[methyl(phenylmethyl)amino]ethyl]oxy)phenyl]pyrazine-2-carboxamide	B
184	3-amino-N-methyl-6-[3-([4-(phenyloxy)phenyl]methyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
185	3-amino-N-methyl-6-[3-([2-(morpholin-4-ylethyl]oxy)phenyl]pyrazine-2-carboxamide	C
186	3-amino-6-[3-([2-((cyclopropylmethyl)(propyl)amino]ethyl]oxy)phenyl]-N-methylpyrazine-2-carboxamide	B
187	3-amino-N-methyl-6-[3-([2-[4-(phenylmethyl)piperidin-1-yl]ethyl]oxy)phenyl]pyrazine-2-carboxamide	B
188	3-amino-N-methyl-6-[3-([2-piperidin-1-ylethyl]oxy)phenyl]pyrazine-2-carboxamide	B
189	3-amino-N-methyl-6-[3-([3-(phenyloxy)phenyl]methyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
190	3-amino-6-(3-([2-(4-ethylpiperazin-1-yl)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide	B
191	3-amino-N-methyl-6-[3-([2-(pyrrolidin-1-ylethyl]oxy)phenyl]pyrazine-2-carboxamide	B
192	3-amino-N-methyl-6-(3-([2-(4-methylpiperazin-1-yl)ethyl]oxy)phenyl)pyrazine-2-carboxamide	B
193	3-amino-6-[3-([2-[4-(2-hydroxyethyl)piperazin-1-yl]ethyl]oxy)phenyl]-N-methylpyrazine-2-carboxamide	B
194	3-amino-6-(3-([2-(3,4-dihydroisoquinolin-2(1H)-yl)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide	B
195	ethyl 4-[2-([3-(5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)phenyl]oxy)ethyl]piperazine-1-carboxylate	C
196	3-amino-N-methyl-6-[3-([3-(3S)-1-(phenylmethyl)pyrrolidin-3-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
197	3-amino-N-methyl-6-[3-([3-(3R)-1-(phenylmethyl)pyrrolidin-3-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
198	3-amino-N-methyl-6-[3-([2-(phenyloxy)ethyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
199	3-amino-N-methyl-6-[3-([2-[4-(phenylmethyl)piperazin-1-yl]ethyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
200	3-amino-N-methyl-6-[3-([1-(phenylmethyl)piperidin-4-yl]methyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	D
201	3-amino-6-(3-([(furan-3-ylmethyl)amino]carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
202	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-2-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
203	3-amino-6-(3-(((furan-2-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
204	3-amino-N-methyl-6-(3-(((4-(phenylmethyl)morpholin-2-yl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	B
205	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	A
206	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	A
207	3-amino-6-(3-(((2,3-dihydro-1-benzofuran-5-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
208	3-amino-6-(3-(((6-fluoro-4H-1,3-benzodioxin-8-yl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
209	3-amino-6-(3-(((2,4-dichlorophenyl)acetyl)amino)phenyl)-N-methylpyrazine-2-carboxamide	A
210	3-amino-6-(3-(((3,4-dichlorophenyl)acetyl)amino)phenyl)-N-methylpyrazine-2-carboxamide	B
211	3-amino-N-methyl-6-(3-(((4-(trifluoromethyl)phenyl)acetyl)amino)phenyl)pyrazine-2-carboxamide	B
212	3-amino-6-(3-(((3,5-dimethylphenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
213	3-amino-6-(3-(((5-chloro-2-methylphenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
214	3-amino-6-(3-(((2,4-dimethylphenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
215	3-amino-6-(3-(((2,3-difluorophenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	C
216	3-amino-N-methyl-6-(3-(((4-(methylthio)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	A
217	3-amino-N-methyl-6-(3-(((2-methylpropyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
218	3-amino-N-methyl-6-(3-(((3-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
219	3-amino-6-(3-(((cyclopropylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
220	3-amino-6-(3-(((cyclopentylamino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	C
221	3-amino-N-methyl-6-(3-(((4-methylphenyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
222	3-amino-N-methyl-6-(3-(((4-(methyloxy)phenyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	B
223	3-amino-6-(3-(((1,3-benzodioxol-5-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B
224	3-amino-6-(3-(((biphenyl-3-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
225	3-amino-N-methyl-6-(3-(((1-methyl-1-phenylethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
226	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
227	3-amino-N-methyl-6-(3-(((4-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	B
228	3-amino-6-[3-(((4-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
229	3-amino-6-[3-(((4-chloro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
230	3-amino-6-[3-(((4-aminophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
231	3-amino-6-[3-(((2-aminophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
232	3-amino-6-[3-(((3-aminophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
233	3-amino-N-methyl-6-pyrimidin-5-ylpyrazine-2-carboxamide	D
234	3-amino-N-methyl-6-[3-(methylsulfonyl)phenyl]pyrazine-2-carboxamide	C
235	3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide	B
236	3-amino-6-(4-hydroxyphenyl)-N-methylpyrazine-2-carboxamide	B
237	3-amino-6-[3-(((4-(aminosulfonyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
238	3-amino-6-[3-(((4-(dimethylamino)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
239	3-amino-6-[3-(((3-bromophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
240	3-amino-6-(4-chloro-3-((phenylmethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
241	methyl 3-(5-amino-6-((cyclopropylamino)carbonyl)pyrazin-2-yl)benzoate	D
242	methyl 3-(5-amino-6-((ethylamino)carbonyl)pyrazin-2-yl)benzoate	C
243	3-amino-6-[3-(((2,3-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
244	3-amino-6-[3-(((4-hydroxy-3-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
245	3-amino-6-[3-(((4-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
246	3-amino-6-[3-(((2-(ethyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
247	3-amino-6-{3-[[[2-chloro-6-(phenyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
248	3-amino-6-{3-[[[2,5-dichlorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
249	3-amino-6-{3-[[[2,6-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	C
250	3-amino-6-{3-[[[2-chloro-6-methylphenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
251	3-amino-N-methyl-6-{3-[[[2,4,5-trifluorophenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide	B
252	3-amino-6-{3-[[[2,2-dimethylpropyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	C
253	3-amino-6-{3-[[[cyclopentylmethyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
254	3-amino-6-{3-[[[3,5-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
255	3-amino-N-methyl-6-{3-[[[4-(methylsulfonyl)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide	B
256	3-amino-6-{3-[[[5-bromo-2-fluorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
257	3-amino-6-{3-[[[3-bromo-4-fluorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
258	3-amino-N-methyl-6-{3-[[[2,4,6-tris(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide	B
259	3-amino-N-methyl-6-{3-[[[3-methylphenyl]methyl]amino]carbonyl]phenyl}pyrazine-2-carboxamide	B
260	3-amino-6-{3-[[[4-bromo-2-fluorophenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
261	3-amino-6-{3-[[[2,5-dimethylphenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
262	3-amino-6-{3-[[[3,5-bis(trifluoromethyl)phenyl]methyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
263	3-amino-6-{3-[[[2,2-diphenylethyl]amino]carbonyl]phenyl}-N-methylpyrazine-2-carboxamide	B
264	3-amino-N-methyl-6-(3-thienyl)pyrazine-2-carboxamide	B
265	3-amino-N-methyl-6-{3-[[[phenylmethyl]amino]sulfonyl]phenyl}pyrazine-2-carboxamide	C
266	5,5'-diamino-N,N'-dimethyl-2,2'-bipyrazine-6,6'-dicarboxamide	D
267	3-amino-6-(1H-indol-5-yl)-N-methylpyrazine-2-carboxamide	B
268	3-amino-N-methyl-6-(2-thienyl)pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
269	3-amino-N-methyl-6-pyridin-4-ylpyrazine-2-carboxamide	C
270	3-amino-N-cyclopropyl-6-(3- (((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
271	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-cyclopropylpyrazine-2-carboxamide	D
272	3-amino-N-ethyl-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
273	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-ethylpyrazine-2-carboxamide	C
274	3-amino-6-[3-(((2,3-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
275	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	A
276	3-amino-6-[3-(((3,5-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
277	3-amino-6-[3-(((5-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
278	3-amino-6-[3-((cyclopropylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	C
279	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
280	3-amino-6-[3-(((4'-fluorobiphenyl-2-yl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
281	3-amino-N-methyl-6-[3-(((2,3,4-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
282	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
283	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
284	3-amino-6-[3-((9H-fluoren-9-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	D
285	3-amino-N-methyl-6-[3-(((2-methylphenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
286	3-amino-6-[3-(((3,4-dimethylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
287	3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
288	3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
289	3-amino-6-[3-(((2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
290	3-amino-N-methyl-6-[3-(((4-(4-methylpiperazin-1-yl)phenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	C
291	3-amino-6-(4-fluoro-3-(((phenylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
292	3-amino-N-methyl-6-[3-(((2,4,6-trichlorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
293	3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
294	3-amino-6-[3-(((2,5-bis(methyloxy)phenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
295	3-amino-6-[3-(((3-fluoro-4-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
296	3-amino-6-[3-(((2-chloro-6-fluoro-3-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
297	3-amino-6-[3-(((2-chloro-3,6-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
298	3-amino-6-[3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
299	3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
300	3-amino-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
301	3-amino-6-[3-(((6-chloro-2-fluoro-3-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
302	3-amino-6-[3-(((4-iodophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	A
303	3-amino-6-[3-(((3-chloro-4-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
304	3-amino-6-[3-(((4-bromophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
305	3-amino-6-[3-(((2-bromophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide	B
306	3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
307	3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
308	3-amino-N-methyl-6-(3-(((phenylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
309	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
310	3-amino-6-[3-(((2,5-difluorophenyl)carbonyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
311	3-amino-N-methyl-6-(3-(((naphthalen-2-ylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
312	3-amino-N-methyl-6-(3-(((phenylmethyl)thio)methyl)phenyl)pyrazine-2-carboxamide	B
313	3-amino-N-methyl-6-(3-(((phenylmethyl)sulfonyl)methyl)phenyl)pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
314	3-amino-N-methyl-6-(3-(((phenylmethyl)oxy)methyl)phenyl)pyrazine-2-carboxamide	B
315	3-amino-N-methyl-6-(3-(((phenylmethyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
316	3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
317	9H-fluoren-9-ylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate	C
318	phenylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate	B
319	ethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate	B
320	3-amino-6-[3-(((3,4-dichlorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
321	1,1-dimethylethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate	C
322	(1R,2S,5R)-5-methyl-2-(1-methylethyl)cyclohexyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate	D
323	3-amino-6-[3-(((4-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
324	3-amino-N-methyl-6-(3-(((4-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide	B
325	3-amino-N-methyl-6-[3-(((2-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
326	3-amino-6-[3-(((3-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
327	3-amino-6-[3-(((2-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
328	3-amino-N-methyl-6-[3-[(2-phenylhydrazino)carbonyl]phenyl]pyrazine-2-carboxamide	A
329	3-amino-N-(2-hydroxyethyl)-6-(3-methylphenyl)pyrazine-2-carboxamide	C
330	3-amino-N-methyl-6-(3-(((phenylacetyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
331	3-amino-6-[3-(((4-chlorophenyl)acetyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
332	3-amino-6-[3-(((biphenyl-4-ylcarbonyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	A
333	3-amino-6-[3-(((2-chloro-5-(trifluoromethyl)phenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
334	N-[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)methyl]-1H-indole-2-carboxamide	B
335	3-amino-6-[3-(((3-hydroxypyridin-2-yl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
336	3-amino-6-[3-(((1H-indol-3-ylacetyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
337	3-amino-6-[3-(((2,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
338	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
339	3-amino-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	A
340	3-amino-6-[3-(((3-iodophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	A
341	3-amino-N-methyl-6-[3-(((2-morpholin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
342	3-amino-N-methyl-6-[3-(((3-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
343	3-amino-N-methyl-6-[3-(((4-morpholin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
344	3-amino-N-methyl-6-[3-(((3-morpholin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
345	3-amino-N-methyl-6-[3-(((4-(1-methylethyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
346	3-amino-N-methyl-6-[3-(((4-(1-methylethyl)phenyl)oxy)acetyl)amino]methyl]phenyl]pyrazine-2-carboxamide	B
347	3-amino-6-[3-(((4-aminophenyl)thio)acetyl)amino]methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
348	3-amino-N-methyl-6-(4-(methyloxy)-3-(((phenylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
349	3-amino-N-methyl-6-(5-(((phenylmethyl)amino)carbonyl]pyridin-3-yl]pyrazine-2-carboxamide	B
350	3-amino-6-[3-(((4-iodophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
351	3-amino-N-methyl-6-[3-(((4-pentylphenyl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
352	3-amino-6-[3-(((1,3-benzodioxol-5-ylcarbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
353	3-amino-6-[3-(((4-(1,1-dimethylethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
354	3-amino-N-methyl-6-[3-(((1-[5-(trifluoromethyl)pyrimidin-2-yl]piperidin-4-yl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
355	3-amino-N-methyl-6-[3-(((2-piperidin-1-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
356	3-amino-N-methyl-6-[3-(((2-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
357	1,1-dimethylethyl (3R)-3-(((3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbonyl)amino)pyrrolidine-1-carboxylate	C
358	3-amino-N-methyl-6-[3-(((3R)-pyrrolidin-3-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide	C

#	Name	Chk1 IC ₅₀
359	3-[[[4-chlorophenyl)methyl]amino]-6-[3-[[[(3R)-1-[(4-chlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
360	3-amino-6-[3-[[[(3R)-1-[(2,4-dichlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
361	N-methyl-3-[[[3-(methyloxy)phenyl]methyl]amino]-6-[3-[[[(3R)-1-[[3-(methyloxy)phenyl]methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
362	3-amino-6-[3-[[[(3R)-1-[(2,6-dichlorophenyl)methyl]pyrrolidin-3-yl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
363	N-methyl-3-[(phenylmethyl)amino]-6-[3-[[[(3R)-1-(phenylmethyl)pyrrolidin-3-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
364	1,1-dimethylethyl 3-[[[3-(5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)phenyl]carbonyl]amino)methyl]pyrrolidine-1-carboxylate	B
365	1,1-dimethylethyl 2-[[[3-(5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)phenyl]carbonyl]amino)methyl]pyrrolidine-1-carboxylate	C
366	3-amino-N-methyl-6-[3-[[[pyrrolidin-3-ylmethyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
367	3-amino-N-methyl-6-[3-[[[pyrrolidin-2-ylmethyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
368	3-amino-6-(2-fluoro-5-[[[phenylmethyl]amino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide	B
369	3-amino-6-[3-[[[1,3-benzodioxol-5-ylacetyl]amino]phenyl]-N-methylpyrazine-2-carboxamide	B
370	3-amino-6-[3-[[[2-chloro-4-fluorophenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
371	3-amino-6-[3-[[[3,4-bis(methyloxy)phenyl]methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
372	3-amino-N-methyl-6-[3-[[[phenylmethyl]amino]phenyl]pyrazine-2-carboxamide	B
373	3-amino-N-methyl-6-[3-[[[2-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
374	3-amino-N-methyl-6-[3-[[[4-(1-methylethyl)phenyl]carbonyl]amino)methyl]phenyl]pyrazine-2-carboxamide	B
375	3-amino-6-[3-[[[4-butylphenyl)methyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
376	3-amino-N-methyl-6-[3-[[[4-(1,2,3-thiadiazol-4-yl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
377	3-amino-N-methyl-6-[3-[[[4-(2-thienyl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
378	3-amino-6-[3-[[[4-(1,1-dimethylethyl)phenyl]carbonyl]amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
379	3-amino-N-methyl-6-[3-[[[4-(1H-pyrazol-1-yl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
380	3-amino-6-[3-[[[3,4-dihydro-2H-1,4-benzoxazin-6-ylmethyl]amino]carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
381	3-amino-6-[3-(((2-amino-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
382	3-amino-6-(3-(((biphenyl-4-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
383	3-amino-6-(3-(((biphenyl-2-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
384	3-amino-6-(3-(((biphenyl-3-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
385	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)pyrrolidin-2-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
386	3-amino-6-(3-(((3,4-dihydro-2H-1,4-benzoxazin-6-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
387	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
388	3-amino-6-[3-(((2,5-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
389	3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
390	3-amino-N-methyl-6-[3-(((2-pyridin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
391	3-amino-N-methyl-6-[3-(((2-pyridin-3-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
392	3-amino-N-methyl-6-[3-(((4-(methylsulfonyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
393	3-amino-6-(3-(((2,2-diphenylethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
394	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)pyrrolidin-3-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
395	3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
396	3-amino-6-[3-(((4-aminophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
397	3-amino-N-methyl-6-[3-(((4-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
398	3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
399	3-amino-6-[3-(((2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
400	3-amino-6-[5-(((2,6-difluorophenyl)methyl)amino)carbonyl]pyridin-3-yl]-N-methylpyrazine-2-carboxamide	B
401	3-amino-6-(5-(((biphenyl-2-ylmethyl)amino)carbonyl)pyridin-3-yl)-N-methylpyrazine-2-carboxamide	B
402	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
403	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
404	3-amino-N-methyl-6-[3-(((2-piperidin-1-ylphenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
405	3-amino-6-[3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
406	3-amino-N-methyl-6-[3-(((6-(trifluoromethyl)pyridin-3-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
407	3-amino-6-[3-(1H-benzimidazol-2-yl)phenyl]-N-methylpyrazine-2-carboxamide	B
408	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
409	3-amino-6-[3-(((3,5-dichlorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
410	3-amino-N-methyl-6-[3-((2-phenylethyl)amino)phenyl]pyrazine-2-carboxamide	B
411	3-amino-6-(5-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]pyridin-3-yl)-N-methylpyrazine-2-carboxamide	B
412	3-amino-N-methyl-6-[3-(((2-(phenylamino)ethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
413	3-amino-6-[3-(((4-ethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide	B
414	3-amino-N-methyl-6-[3-(((4-pyridin-3-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
415	3-amino-6-[3-((2,3-dihydro-1H-inden-2-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	A
416	3-amino-6-[3-(1,3-dihydro-2H-isolindol-2-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide	B
417	3-amino-6-[3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
418	3-amino-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	A
419	3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	A
420	3-amino-N-methyl-6-[3-(((1R)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
421	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
422	3-amino-N-methyl-6-[3-(((4-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	A
423	3-amino-N-methyl-6-[3-(((3-(1,2,3-thiadiazol-4-yl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
424	3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
425	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
426	3-amino-N-methyl-6-[3-(((3-(trifluoromethyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
427	3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
428	3-amino-6-[3-(((1R)-2-hydroxy-1-phenylethyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
429	methyl (2S)-{[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)methyl]amino}(phenyl)ethanoate	C
430	3-amino-6-(3-{[2-(2,5-difluorophenyl)hydrazino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide	B
431	3-amino-N-methyl-6-(3-{[2-(2,3,5,6-tetrafluoro-4-methylphenyl)hydrazino]carbonyl}phenyl)pyrazine-2-carboxamide	B
432	3-amino-6-(3-{[2-(2-fluorophenyl)hydrazino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide	B
433	3-amino-6-(3-{[2-(2,4-difluorophenyl)hydrazino]carbonyl}phenyl)-N-methylpyrazine-2-carboxamide	B
434	3-amino-N-methyl-6-(3-{[2-(2,3,5,6-tetrafluorophenyl)hydrazino]carbonyl}phenyl)pyrazine-2-carboxamide	B
435	3-amino-N-methyl-6-[3-(((5-methylpyrazin-2-yl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
436	3-amino-N-methyl-6-(3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	A
437	3-amino-N-methyl-6-[3-(((3S)-1-(phenylmethyl)pyrrolidin-3-yl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
438	3-amino-N-methyl-6-[3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
439	3-amino-6-(3-{[ethyl(phenylmethyl)amino]methyl}phenyl)-N-methylpyrazine-2-carboxamide	B
440	3-amino-N-methyl-6-[3-(((1S,2S)-2-phenylcyclopropyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
441	3-Amino-6-(3-benzylcarbonyl-phenyl)-pyrazine-2-carboxylic acid methylamide	B
442	3-amino-N-methyl-6-[3-(4-phenyl-1H-imidazol-2-yl)phenyl]pyrazine-2-carboxamide	B
443	3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
444	3-amino-6-[3-(((5-bromo-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
445	3-amino-N-methyl-6-(3-(((2R)-1,2,3,4-tetrahydronaphthalen-2-ylamino)methyl)phenyl)pyrazine-2-carboxamide	A
446	3-amino-N-methyl-6-(3-(((1-methyl-1-phenylethyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
447	3-amino-6-[3-[(9H-fluoren-9-ylamino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	C
448	3-amino-N-methyl-6-[3-[(naphthalen-1-ylamino)methyl]phenyl]pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
449	3-amino-6-[3-(((3-fluorobiphenyl-4-yl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
450	3-amino-6-[3-(((2-fluoro-4-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
451	3-amino-6-[3-(((2-fluoro-4-furan-2-ylphenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
452	3-amino-N-methyl-6-[3-((naphthalen-2-ylamino)methyl)phenyl]pyrazine-2-carboxamide	B
453	3-amino-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
454	3-amino-N-methyl-6-[3-(((2S)-1,2,3,4-tetrahydronaphthalen-2-ylamino)methyl)phenyl]pyrazine-2-carboxamide	A
455	methyl 3-amino-6-[3-(((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxylate	B
456	3-amino-6-[3-(((5-bromo-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
457	3-amino-6-[3-(((2-fluoro-5-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
458	3-amino-6-[3-(((4-ethenyl-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
459	3-amino-6-[3-(((4,7-difluoro-3-methyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
460	3-amino-N-methyl-6-[3-(((1S,2R)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
461	3-amino-6-[3-(((5-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
462	3-amino-6-[3-(((2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
463	3-amino-6-[3-(((2R)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
464	3-amino-6-[3-(((5-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
465	3-amino-6-[3-(((2S)-5-bromo-2,3-dihydro-1H-inden-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
466	3-amino-6-[3-(((4-fluoro-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	A
467	3-amino-6-[3-(((4-fluoro-1-hydroxy-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
468	3-amino-N-methyl-6-[3-(((1R,2S)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
469	3-amino-6-[3-(((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
470	3-amino-6-[3-(((6-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
471	3-amino-N-methyl-6-[3-(((3-oxo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
472	3-amino-6-(3-(((5-furan-2-yl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	A
473	3-amino-6-(3-(((4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
474	3-amino-6-(3-(((4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
475	3-amino-6-(3-(((6-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	A
476	3-amino-6-(3-(((5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
477	3-amino-6-(3-(((5-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	A
478	3-amino-N-methyl-6-[3-(((5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	A
479	3-amino-6-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
480	3-amino-6-(3-(((5-furan-3-yl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	A
481	3-amino-N-methyl-6-[3-(((5-(3-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	A
482	3-amino-6-[3-(((1S)-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	A
483	3-amino-6-[3-(((1S)-5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide	B
484	3-amino-6-(3-(((7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
485	3-amino-6-(3-(((6-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
486	3-amino-6-(3-(((4,6-dichloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	B
487	3-amino-6-(3-(((4-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	A
488	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-ethylpyrazine-2-carboxamide	B
489	3-amino-6-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-ethylpyrazine-2-carboxamide	B
490	3-amino-N-methyl-6-[3-(((1S)-5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide	A
491	3-amino-N-methyl-6-[3-(((5-phenyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
492	3-amino-6-(3-(((4-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide	A
493	3-amino-N-methyl-6-[3-(((4-phenyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	A
494	3-amino-N-methyl-6-[3-(((4-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
495	3-amino-N-methyl-6-[3-(((6-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
496	3-amino-N-methyl-6-[3-(((5-(4-methyl-2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	A
497	3-amino-N-methyl-6-[3-(((2R)-6-phenyl-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
498	3-amino-N-methyl-6-naphthalen-2-ylpyrazine-2-carboxamide	B
499	3-amino-N-methyl-6-[3-(((1S,2S)-2-((phenylmethyl)oxy)cyclopentyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
500	3-amino-6-[3-((cyclopentylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
501	3-amino-6-[3-(((6-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
502	3-amino-6-[3-(((7-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
503	3-amino-N-methyl-6-quinolin-3-ylpyrazine-2-carboxamide	C
504	3-amino-6-[3-(1H-imidazol-1-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide	B
505	3-amino-N-methyl-6-[3-(((5-((2-morpholin-4-ylethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
506	3-amino-N-methyl-6-[3-(((1R,2R)-2-((phenylmethyl)oxy)cyclopentyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
507	3-amino-6-[3-((2,3-dihydro-1-benzofuran-3-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
508	3-amino-6-[3-(((5-((cyanomethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
509	2-amino-5-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]-N-methylpyridine-3-carboxamide	B
510	3-amino-N-methyl-6-[3-(((5-(4-methylphenyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
511	3-amino-6-[3-(((2-(dimethylamino)-1-phenylethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
512	3-amino-N-methyl-6-[3-(((5-(1H-1,2,4-triazol-1-yl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
513	3-amino-6-[3-(((5-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	B
514	3-amino-N-methyl-6-[3-(((4-((2-morpholin-4-ylethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide	B
515	3-amino-6-[3-(((4-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide	A
516	3-amino-N-methyl-6-[1-((phenylmethyl)amino)isoquinolin-7-yl]pyrazine-2-carboxamide	C
517	3-amino-N-methyl-6-[3-(((2-morpholin-4-yl-1-phenylethyl)amino)methyl)phenyl]pyrazine-2-carboxamide	C

#	Name	Chk1 IC ₅₀
518	3-amino-N-methyl-6-(3-(((1-phenyl-2-pyrrolidin-1-ylethyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
519	2-amino-5-(3-(((6-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide	B
520	2-amino-5-(3-(((4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide	B
521	2-amino-5-(3-(((7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide	C
522	2-amino-5-(3-(((4-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide	B
523	2-amino-5-(3-(((5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide	B
524	2-amino-5-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide	B
525	3-amino-6-(1-(((1S)-2,3-dihydro-1H-inden-1-ylamino)isoquinolin-7-yl))-N-methylpyrazine-2-carboxamide	B
526	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)-2,3-dihydro-1H-inden-5-yl))-N-methylpyrazine-2-carboxamide	A
527	3-amino-6-(3-(((6-(3-hydroxypropyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl))-N-methylpyrazine-2-carboxamide	A
528	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(1,3-oxazol-5-yl)pyrazin-2-amine	C
529	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carbohydrazide	B
530	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	B
531	1-[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazin-2-yl]ethanone	B
532	3-amino-6-(3-(((2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl))-N-(naphthalen-2-ylmethyl)pyrazine-2-carboxamide	D
533	3-amino-N-cyclohexyl-6-(3-(((2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	C
534	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(pyridin-4-ylmethyl)pyrazine-2-carboxamide	C
535	{3-[5-amino-6-(4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol	C
536	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(4H-1,2,4-triazol-3-yl)pyrazin-2-amine	B
537	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2,2,2-trifluoroethyl)pyrazine-2-carboxamide	C
538	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(5-hydroxypentyl)pyrazine-2-carboxamide	C
539	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(4-hydroxybutyl)pyrazine-2-carboxamide	B
540	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-pyrrolidin-1-ylethyl)pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
541	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-piperidin-1-ylethyl)pyrazine-2-carboxamide	C
542	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-morpholin-4-ylethyl)pyrazine-2-carboxamide	C
543	3-amino-N-(cyclopropylmethyl)-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	B
544	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(phenylmethyl)pyrazine-2-carboxamide	C
545	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-phenylethyl)pyrazine-2-carboxamide	C
546	[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazin-2-yl](phenyl)methanone	B
547	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-hydroxyethyl)pyrazine-2-carboxamide	B
548	3-amino-N-cyclopropyl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	B
549	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine	C
550	{3-[5-amino-6-(5-methyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol	C
551	{3-[5-amino-6-(5-phenyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol	B
552	(3-{5-amino-6-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol	C
553	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N'-methylpyrazine-2-carbohydrazide	B
554	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N,N-dimethylpyrazine-2-carboxamide	B
555	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine	B
556	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(dimethylamino)ethyl]pyrazine-2-carboxamide	B
557	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-phenyl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	A
558	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide	A
559	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(methylamino)ethyl]pyrazine-2-carboxamide	A
560	3-{5-amino-6-[(3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
561	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(3-hydroxypropyl)pyrazine-2-carboxamide	B
562	1,1-dimethylethyl 4-(((3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate	D
563	1,1-dimethylethyl 4-(((3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)methylpiperidine-1-carboxylate	D

#	Name	Chk1 IC ₅₀
564	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-piperidin-4-ylpyrazine-2-carboxamide	B
565	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-(piperidin-4-ylmethyl)pyrazine-2-carboxamide	B
566	3-[5-amino-6-(morpholin-4-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
567	1,1-dimethylethyl 3-(((3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	D
568	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-pyrrolidin-3-ylpyrazine-2-carboxamide	B
569	(3-{5-amino-6-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol	C
570	{3-[5-amino-6-(5-furan-2-yl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl)methanol	B
571	[3-(5-amino-6-{5-[3-(methyloxy)phenyl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol	D
572	2-((3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl)carbonyl)-N-phenylhydrazinecarboxamide	B
573	3-[5-amino-6-[(4-methylpiperazin-1-yl)carbonyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
574	3-(5-amino-6-[(4-(4-fluorophenyl)piperazin-1-yl)carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
575	3-(5-amino-6-[(4-(phenylmethyl)piperazin-1-yl)carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
576	methyl N-[[3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]glycinate	C
577	N-[[3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]glycine	D
578	3-{5-amino-6-[(3,5-difluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
579	3-[5-amino-6-(biphenyl-4-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
580	{3-[5-amino-6-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl)methanol	B
581	(3-{5-amino-6-[3-(4-chlorophenyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol	D
582	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine	B
583	(3-{5-amino-6-[3-(2-thienyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl}phenyl)methanol	B
584	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[3-(3-(methyloxy)phenyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-amine	B
585	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[3-(dimethylamino)propyl]pyrazine-2-carboxamide	C
586	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[3-(1H-imidazol-1-yl)propyl]pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
587	3-{5-amino-6-[(4-chloro-3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
588	3-(5-amino-6-[[2,4-bis(methoxy)phenyl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
589	3-(5-amino-6-[[4-(dimethylamino)phenyl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
590	methyl 1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]-L-prolinate	D
591	methyl 1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]piperidine-3-carboxylate	D
592	1,1-dimethylethyl 4-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]piperazine-1-carboxylate	D
593	3-[5-amino-6-(piperazin-1-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
594	N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]glutamic acid	C
595	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-{5-[[methoxy)methyl]-4H-1,2,4-triazol-3-yl}pyrazin-2-amine	C
596	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	A
597	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[3-(methylamino)propyl]pyrazine-2-carboxamide	A
598	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-(3-pyrrolidin-1-ylpropyl)pyrazine-2-carboxamide	C
599	3-(5-amino-6-[(1E)-N-[4-(methoxy)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	C
600	3-(5-amino-6-[(1E)-N-[4-(1-methylethyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	C
601	3-{5-amino-6-[(1E)-N-1,3-benzothiazol-2-ylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
602	3-{5-amino-6-[(1E)-N-(4-methylphenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
603	3-{5-amino-6-[(1E)-N-(4-chlorophenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
604	3-{5-amino-6-[(1E)-N-methyl-N-phenylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
605	3-{5-amino-6-[(1E)-N-(2-hydroxyethyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	B
606	{3-[5-amino-6-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol	A
607	{3-[5-amino-6-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol	C
608	3-[5-amino-6-(2-aminopyrimidin-4-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
609	3-{5-amino-6-[2-(dimethylamino)pyrimidin-4-yl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D

#	Name	Chk1 IC ₅₀
610	ethyl N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]-L-tyrosinate	D
611	N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]-D-tyrosine	D
612	1,1-dimethylethyl [3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]amino]propyl]carbamate	D
613	3-amino-N-(3-aminopropyl)-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
614	1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]piperidine-3-carboxylic	D
615	1,1-dimethylethyl [2-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]amino]ethyl]carbamate	D
616	3-amino-N-(2-aminoethyl)-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	B
617	1,1-dimethylethyl (1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl)carbonyl]piperidin-4-yl]carbamate	D
618	3-{5-amino-6-[(4-aminopiperidin-1-yl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
619	3-{5-amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
620	3-{5-amino-6-[imino(morpholin-4-ylamino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
621	3-(5-amino-6-[imino[(4-methylpiperazin-1-yl)amino]methyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
622	3-{5-amino-6-[imino(piperidin-1-ylamino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
623	3-{5-amino-6-[(azepan-1-ylamino)(imino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
624	3-{5-amino-6-[imino(((2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl)amino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
625	{3-[5-amino-6-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)]pyrazin-2-yl}phenyl]methanol	B
626	3-{5-amino-6-[(1E)-N-morpholin-4-ylethanimidoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
627	4-((2E)-2-{1-[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]ethylidene)hydrazino]benzoic acid	C
628	ethyl ((2E)-2-{1-[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]ethylidene)hydrazino]acetate	D
629	3-{5-amino-6-[(1E)-N,N-dimethylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	B
630	3-(5-amino-6-[(1E)-N-[4-(methylsulfonyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
631	3-{5-amino-6-[(1E)-N-(4-cyanophenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D

#	Name	Chk1 IC ₅₀
632	3-{5-amino-6-[(1E)-N-pyridin-2-ylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
633	3-(5-amino-6-[(1E)-N-[amino(imino)methyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	C
634	3-[5-amino-6-[(1E)-N-[4-[(trifluoromethyl)oxy]phenyl]ethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
635	3-(5-Amino-6-[1-[(4-nitro-phenyl)-hydrazono]-ethyl]-pyrazin-2-yl)-N-benzylbenzamide	D
636	3-(5-amino-6-[(1E)-N-[4-(trifluoromethyl)pyrimidin-2-yl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	C
637	3-{5-amino-6-[(1E)-N-1H-1,2,3-benzotriazol-1-ylethanimidoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
638	3-{5-amino-6-[(1E)-N-methylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
639	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	B
640	3-(5-amino-6-[(1E)-N-[4-(trifluoromethyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
641	3-{5-amino-6-[(1E)-N-phenylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
642	3-{5-amino-6-[(1E)-N-(4-methylpiperazin-1-yl)ethanimidoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	B
643	{3-[5-amino-6-(3-pyrrolidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol	B
644	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
645	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
646	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(furan-2-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide	C
647	N-[3-(acetilamino)propyl]-3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	C
648	3-amino-N-[3-[(furan-2-ylcarbonyl)amino]propyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide	C
649	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(pyridin-4-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide	D
650	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(pyridin-4-ylcarbonyl)amino]propyl]pyrazine-2-carboxamide	C
651	N-[3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]propyl]quinoxaline-2-carboxamide	D
652	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[[[6-(trifluoromethyl)pyridin-3-yl]carbonyl]amino]propyl]pyrazine-2-carboxamide	D
653	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(quinolin-8-ylsulfonyl)amino]propyl]pyrazine-2-carboxamide	D
654	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(pyridin-2-ylsulfonyl)amino]propyl]pyrazine-2-carboxamide	C

#	Name	Chk1 IC ₅₀
655	3-amino-N-[3-(((3,5-dimethylisoxazol-4-yl)amino)carbonyl)amino]propyl]-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
656	3-[5-amino-6-(2-methylpyrimidin-4-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
657	3-[5-amino-6-[2-(methylamino)pyrimidin-4-yl]pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
658	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-[2-[(pyridin-4-ylcarbonyl)amino]ethyl]pyrazine-2-carboxamide	C
659	3-amino-N-[2-[(furan-2-ylcarbonyl)amino]ethyl]-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	D
660	N,N'-cyclohexane-1,2-diylbis[3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide]	D
661	N-[2-(acetylamino)ethyl]-3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
662	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-[2-[(quinolin-8-ylsulfonyl)amino]ethyl]pyrazine-2-carboxamide	D
663	N-[2-(((3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)ethyl]quinoxaline-2-carboxamide	D
664	3-amino-N-[2-[(2-chloropyridin-3-yl)carbonyl]amino]ethyl]-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
665	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-[2-[(pyridin-2-ylsulfonyl)amino]ethyl]pyrazine-2-carboxamide	D
666	3-amino-N-[2-(((3,5-dimethylisoxazol-4-yl)amino)carbonyl)amino]ethyl]-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	C
667	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide	B
668	3-[5-amino-6-(imino{2-[4-(trifluoromethyl)pyrimidin-2-yl]hydrazino)methyl}pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
669	3-[5-amino-6-[[2-(1,3-benzothiazol-2-yl)hydrazino](imino)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
670	3-[5-amino-6-(1,5-diphenyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
671	3-[5-amino-6-(1,5-dimethyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
672	3-[5-amino-6-(1-methyl-5-phenyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
673	3-[5-amino-6-((1E)-N-((2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl)ethanimidoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	B
674	3-[5-amino-6-[(1E)-N-azepan-1-ylethanimidoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide	B
675	3-(5-amino-6-((E)-[(phenylmethyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
676	3-[5-amino-6-((E)-[amino(imino)methyl]hydrazono)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
677	3-(5-amino-6-((E)-[(2-hydroxyethyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide	C

#	Name	Chk1 IC ₅₀
678	3-{5-amino-6-[(E)-(pyridin-2-ylhydrazono)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
679	3-(5-amino-6-[(E)-{(4-cyanophenyl)hydrazono}methyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
680	3-(5-amino-6-[(E)-{(4-methylphenyl)hydrazono}methyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide	D
681	3-{5-amino-6-[(E)-(hydroxyimino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	C
682	{3-[5-amino-6-(3-pyridin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol	C
683	5-(3-[[{(1R)-2,3-dihydro-1H-inden-1-ylamino}methyl]phenyl]-3-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	A
684	3-amino-N-[3-[[{4-(dimethylamino)phenyl}carbonyl]amino]propyl]-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazine-2-carboxamide	D
685	3-amino-6-(3-[[{(1S)-2,3-dihydro-1H-inden-1-ylamino}methyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
686	1,1-dimethylethyl 4-[(3S)-3-[[{3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazin-2-yl}carbonyl]amino]pyrrolidin-1-yl]piperidine-1-carboxylate	C
687	3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]-N-[(3S)-1-piperidin-4-ylpyrrolidin-3-yl]pyrazine-2-carboxamide	C
688	1,1-dimethylethyl (2R)-2-[[{(3S)-3-[[{3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazin-2-yl}carbonyl]amino]pyrrolidin-1-yl]methyl]pyrrolidine-1-carboxylate	D
689	3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]-N-[(3S)-1-[(2R)-pyrrolidin-2-ylmethyl]pyrrolidin-3-yl]pyrazine-2-carboxamide	C
690	3-amino-N-[(3S)-1-[[{4-(dimethylamino)phenyl}methyl]pyrrolidin-3-yl]-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazine-2-carboxamide	C
691	3-amino-N-{2-[[{4-(dimethylamino)phenyl}amino]carbonyl]amino}ethyl]-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazine-2-carboxamide	D
692	3-amino-N-{2-[[{4-(dimethylamino)phenyl}carbonyl]amino}ethyl]-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazine-2-carboxamide	D
693	3-{5-amino-6-[imino(2-pyridin-2-ylhydrazino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide	D
694	3-[5-amino-6-(morpholin-4-ylmethyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
695	ethyl 1-[[{3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazin-2-yl]methyl]piperidine-4-carboxylate	D
696	3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]-N-[(3S)-1-(pyridin-2-ylmethyl)pyrrolidin-3-yl]pyrazine-2-carboxamide	B
697	methyl N-[[{3-amino-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazin-2-yl]carbonyl]threoninate	D
698	{3-[5-amino-6-(3-piperidin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol	D
699	3-(ethylamino)-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazine-2-carboxamide	D
700	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-(3-[[{(phenylmethyl)amino}carbonyl]phenyl]pyrazine-2-carboxamide	B

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701	3-[5-amino-6-(1-methyl-5-piperidin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	D
702	3-[5-amino-6-(5-methyl-1-pyridin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
703	{3-[5-amino-6-(3-morpholin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol	B
704	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	A
705	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-((3R)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	A
706	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-((3S)-1-ethylpyrrolidin-3-yl)pyrazine-2-carboxamide	B
707	6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-3-(ethylamino)-N-((3S)-1-ethylpyrrolidin-3-yl)pyrazine-2-carboxamide	D
708	3-amino-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide	B
709	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	D
710	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((1R)-1-phenylethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	D
711	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(3,4-dihydroisoquinolin-2(1H)-ylcarbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	D
712	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-((2-phenylethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	D
713	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((1S)-1-phenylethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	C
714	3-amino-6-[3-(3,4-dihydroisoquinolin-2(1H)-ylcarbonyl)phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide	B
715	3-amino-6-(3-(((2-phenylethyl)amino)carbonyl)phenyl)-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide	B
716	3-amino-6-[3-(((1R)-1-phenylethyl)amino)carbonyl]phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide	C
717	3-amino-6-[3-(((1S)-1-phenylethyl)amino)carbonyl]phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide	B
718	3-(5-amino-6-[(2-ethylhydrazino)(imino)methyl]pyrazin-2-yl)-N-(2,3-dihydro-1H-inden-1-yl)benzamide	C
719	3-(5-amino-6-[imino(2-methylhydrazino)methyl]pyrazin-2-yl)-N-((1S)-2,3-dihydro-1H-inden-1-yl)benzamide	C
720	3-(5-amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl)-N-((1S)-2,3-dihydro-1H-inden-1-yl)benzamide	D
721	phenylmethyl 3-(3-[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl)piperidine-1-carboxylate	D
722	phenylmethyl 3-(3-[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl)piperidine-1-carboxylate	D

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723	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine	A
724	[3-(5-amino-6-(3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl)pyrazin-2-yl)phenyl]methanol	B
725	[3-(5-amino-6-(3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl)pyrazin-2-yl)phenyl]methanol	B
726	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
727	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	C
728	1,1-dimethylethyl 3-(((3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate	D
729	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)-N-piperidin-3-ylpyrazine-2-carboxamide	A
730	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)-N-(1-ethylpiperidin-3-yl)pyrazine-2-carboxamide	B
731	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	B
732	(3-{5-amino-6-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]}pyrazin-2-yl)phenyl]methanol	C
733	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine	B
734	3-[5-amino-6-(1-ethyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]benzamide	B
735	3-[5-amino-6-(1-methyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	B
736	(3-{5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]}pyrazin-2-yl)phenyl]methanol	C
737	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-amine	B
738	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((4-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	B
739	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((4-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	B
740	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	C
741	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((4-(methylthio)phenyl)methyl)amino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	C
742	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate	B
743	3-amino-6-[3-(((4-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
744	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-[3-(((4-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A

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745	3-amino-6-[3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
746	3-amino-6-[3-[(2-chloro-6-fluorophenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
747	3-amino-6-[3-[(4-(methylthio)phenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
748	3-[5-amino-6-[imino(2-pyridin-4-ylhydrazino)methyl]pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]benzamide	C
749	phenylmethyl (3R)-3-[3-[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate	D
750	3-amino-6-(3-[(1S)-1-[(1S)-2,3-dihydro-1H-inden-1-ylamino]ethyl]phenyl)-N-methylpyrazine-2-carboxamide	B
751	3-amino-6-[3-[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide	B
752	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)pyrazin-2-yl]carbonyl]amino)methyl]pyrrolidine-1-carboxylate	D
753	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide	A
754	phenylmethyl (3R)-3-[3-[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate	C
755	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	A
756	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[(4-chlorophenyl)methyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate	B
757	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(pyridin-2-ylmethyl)amino]carbonyl]phenyl)pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate	C
758	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(pyridin-3-ylmethyl)amino]carbonyl]phenyl)pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate	C
759	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[(3,4-dihydro-2H-chromen-4-ylamino)carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate	C
760	3-amino-6-[3-[(4-chlorophenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
761	3-amino-6-(3-[(pyridin-2-ylmethyl)amino]carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
762	3-amino-6-(3-[(pyridin-3-ylmethyl)amino]carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
763	dimethyl 5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzene-1,3-dicarboxylate	C
764	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
765	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[(3R)-piperidin-3-yl]pyrazine-2-carboxamide	A

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766	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-piperidin-3-ylpyrazine-2-carboxamide	A
767	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide	B
768	3-amino-N-azepan-4-yl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazine-2-carboxamide	B
769	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methylpyrrolidine-1-carboxylate	D
770	1,1-dimethylethyl 4-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate	D
771	3-[5-amino-6-(5-[2-[(phenylmethyl)oxy]ethyl)-1H-1,2,4-triazol-3-yl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	C
772	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)-4-fluorophenyl)-N-methylpyrazine-2-carboxamide	B
773	3-[5-amino-6-(1-phenyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	B
774	3-[5-amino-6-[amino(imino)methyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	C
775	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]benzamide	A
776	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate	B
777	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate	A
778	3-amino-6-[3-((3,4-dihydro-2H-chromen-4-ylamino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
779	3-amino-6-[3-((2,3-dihydro-1-benzofuran-3-ylamino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
780	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide	A
781	1,1-dimethylethyl 4-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate	B
782	3-amino-N-azepan-4-yl-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
783	3-amino-N-azepan-4-yl-6-[3-[(3aR,8aS)-8,8a-dihydro-3aH-indeno[1,2-d][1,3]oxazol-2-yl]phenyl]pyrazine-2-carboxamide	B
784	1,1-dimethylethyl (3R)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)methylpyrrolidine-1-carboxylate	D
785	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide	A
786	1,1-dimethylethyl (3R)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methylpyrrolidine-1-carboxylate	D

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787	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide	A
788	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)-5-fluorophenyl)-N-methylpyrazine-2-carboxamide	B
789	3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-5-[(methoxy)carbonyl]benzoic acid	C
790	methyl 3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-5-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)benzoate	B
791	5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-N'-pyrrolidin-3-ylbenzene-1,3-dicarboxamide	B
792	3-[5-amino-6-(5-phenyl-1-pyridin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide	C
793	3-[5-amino-6-(5-methyl-1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	C
794	1,1-dimethylethyl (3R)-3-[[[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate	C
795	1,1-dimethylethyl (3R)-3-[[[3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl)phenyl]pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate	D
796	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
797	5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-N'-[(3R)-pyrrolidin-3-yl]benzene-1,3-dicarboxamide	B
798	(3-[5-amino-6-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl]phenyl)methanol	B
799	3-[5-amino-6-(1,5-dimethyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	D
800	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(1-methyl-1-phenylethyl)amino]carbonyl)phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate	D
801	3-amino-6-(3-[[[1-methyl-1-phenylethyl]amino]carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
802	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)sulfonyl)phenyl)pyrazine-2-carboxamide	C
803	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)sulfonyl)phenyl)-N-piperidin-3-ylpyrazine-2-carboxamide	B
804	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-[(3S)-1-methylpyrrolidin-3-yl]pyrazine-2-carboxamide	B
805	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl)phenyl]-N-[(3S)-1-methylpyrrolidin-3-yl]pyrazine-2-carboxamide	B
806	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)methyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
807	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
808	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-(3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
809	N-[3-(5-amino-6-(5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl)pyrazin-2-yl)phenyl]-2-(2,4-dichlorophenyl)acetamide	A
810	1,1-dimethylethyl (3S)-3-([(3-amino-6-(3-[(1R)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl)phenyl)pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate	D
811	3-amino-6-(3-hydroxyphenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
812	3-amino-6-(3-[(1R)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl)phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	B
813	3-[5-amino-6-(1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide	C
814	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-[3-([(3,4,5-trifluorophenyl)methyl]amino)carbonyl)phenyl]pyrazine-2-carboxamide	A
815	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-[3-([(2,3,6-trifluorophenyl)methyl]amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
816	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-[3-([(2,3,5-trifluorophenyl)methyl]amino)carbonyl)phenyl]pyrazine-2-carboxamide	B
817	3-amino-6-[3-([(3-bromo-4-fluorophenyl)methyl]amino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
818	3-[5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl]-N-[(2-chloro-6-fluorophenyl)methyl]benzamide	C
819	3-amino-N-1-azabicyclo[2.2.2]oct-3-yl-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl]pyrazine-2-carboxamide	B
820	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(4-chloro-2-fluorophenyl)methyl]benzamide	A
821	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2-chlorophenyl)methyl]benzamide	A
822	1,1-dimethylethyl (3S)-3-([(3-amino-6-[3-([(2,4-dichlorophenyl)methyl]amino)carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate	C
823	1,1-dimethylethyl (3S)-3-([(3-amino-6-[3-([(3,4-dichlorophenyl)methyl]amino)carbonyl)phenyl]pyrazin-2-yl]carbonyl)amino]pyrrolidine-1-carboxylate	C
824	5-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenyl)-3-(4-piperidin-3-yl-1,3-thiazol-2-yl)pyrazin-2-amine	D
825	3-amino-6-[3-([(2,4-dichlorophenyl)methyl]amino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
826	3-amino-6-[3-([(3,4-dichlorophenyl)methyl]amino)carbonyl)phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide	A
827	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2,6-difluorophenyl)methyl]benzamide	B
828	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2-chloro-6-fluorophenyl)methyl]benzamide	B
829	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2,5-dichlorophenyl)methyl]benzamide	B
830	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(3,4-dichlorophenyl)methyl]benzamide	A

#	Name	Chk1 IC ₅₀
831	3-amino-6-[3-[[[4-(methoxy)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
832	3-amino-6-[3-[[[2,4-dichlorophenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
833	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[[[1,1-dimethylethyl]oxy]carbonyl]amino]methyl]phenyl]pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate	B
834	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(2-fluorophenyl)methyl]benzamide	A
835	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(3-fluorophenyl)methyl]benzamide	A
836	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(4-fluorophenyl)methyl]benzamide	A
837	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(3-chlorophenyl)methyl]benzamide	A
838	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(4-chlorophenyl)methyl]benzamide	A
839	3-amino-6-[3-(aminomethyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
840	3-amino-6-(3-hydroxyphenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
841	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate	B
842	3-amino-6-[3-[[[4-fluorophenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
843	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
844	3-amino-6-[3-[[[biphenyl-4-ylmethyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
845	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-(1,2,3-thiadiazol-5-yl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
846	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[2-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
847	3-amino-6-[3-[[[biphenyl-2-ylmethyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
848	3-amino-6-[3-[[[2,4-difluorophenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
849	3-amino-6-[3-[[[4-chloro-2-fluorophenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
850	3-amino-6-[3-[[[4-fluoro-3-(trifluoromethyl)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
851	3-amino-6-[3-[[[naphthalen-2-ylmethyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
852	3-amino-6-[3-[[[4-(dimethylamino)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
853	3-amino-6-(3-(((1,3-benzodioxol-5-ylmethyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
854	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((4-(2-thienyl)phenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	A
855	3-amino-6-(3-(((2-(methyloxy)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
856	3-amino-6-(3-(((2,4-bis(methyloxy)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
857	3-amino-6-(3-(((4-chloro-2-methylphenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
858	3-amino-6-(3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
859	3-amino-6-(3-(((4-(methylsulfonyl)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
860	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
861	3-amino-6-(3-(((4-chlorophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
862	3-amino-6-(3-(((5-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
863	3-amino-N-[(3R)-1-azabicyclo[2.2.2]oct-3-yl]-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide	B
864	3-amino-6-(3-(((4-(methylthio)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
865	3-(5-amino-6-(3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl)pyrazin-2-yl)-N-[(2,4-difluorophenyl)methyl]benzamide	A
866	3-amino-6-(3-(((1,2-diphenylethyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
867	3-(5-amino-6-(3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl)pyrazin-2-yl)-N-[(3,4-difluorophenyl)methyl]benzamide	A
868	3-amino-6-(3-(((3,5-difluorophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
869	methyl 4-((((3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl)pyrazin-2-yl)phenyl)carbonyl)amino)methyl]benzoate	A
870	3-amino-6-(3-(((4-bromophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
871	3-amino-6-(3-(((4-bromo-2-fluorophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
872	3-amino-6-(3-(((2-bromophenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
873	3-amino-6-(3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
874	3-amino-6-(3-(((2-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
875	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3R)-pyrrolidin-3-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
876	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3R)-pyrrolidin-3-ylamino)carbonyl)phenyl]pyrazine-2-carboxamide	A
877	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2-piperidin-1-ylethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	A
878	3-amino-6-[3-(((2-phenylethyl)amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
879	3-amino-6-[3-(((2-(methyloxy)ethyl)amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	B
880	3-amino-6-[3-(((4-iodophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
881	3-amino-6-[3-(((2-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
882	3-amino-6-[3-(((4-fluorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
883	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(trifluoromethyl)oxy)phenyl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	A
884	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
885	3-amino-6-[3-(((4-chlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
886	3-amino-6-[3-(((2,6-dichlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
887	3-amino-6-[3-(((pentafluorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
888	3-amino-6-[3-(((2-chloro-4-fluorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
889	3-amino-6-[3-(((4-(methylthio)phenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
890	3-amino-6-[3-(((2,4-dichlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
891	3-amino-6-[3-(((4-(4-chlorophenyl)-2-thienyl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
892	3-amino-6-[3-(((4-bromophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
893	3-amino-6-[3-(((naphthalen-2-ylcarbonyl)amino)methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
894	3-amino-6-[3-(((2-(4-phenylpiperazin-1-yl)ethyl)oxy)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
895	N-[(3-{5-amino-6-[(3R)-piperidin-3-ylacetyl]pyrazin-2-yl}phenyl)methyl]-N'-(4-bromophenyl)urea	A
896	N-[(3-{5-amino-6-[(3R)-piperidin-3-ylacetyl]pyrazin-2-yl}phenyl)methyl]-N'-naphthalen-2-ylurea	A
897	3-amino-6-[3-(((2,4-difluorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
898	3-amino-6-(3-(((2-fluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
899	3-amino-6-(3-(((5-[2-chloro-5-(trifluoromethyl)phenyl]furan-2-yl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
900	3-amino-6-[3-(((4-(dimethylamino)naphthalen-1-yl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
901	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1,2,3-thiadiazol-5-yl)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide	A
902	3-amino-6-(3-(((1,3-dimethyl-1H-pyrazol-5-yl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
903	3-amino-6-(3-(((4-(dimethylamino)phenyl)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
904	3-amino-6-(3-(((2-hydroxyethyl)oxy)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
905	3-amino-6-(3-(((2,3-dihydro-1-benzofuran-5-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
906	3-amino-6-(3-(((3,4-dihydro-2H-1,5-benzodioxepin-7-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
907	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((quinolin-7-ylsulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide	B
908	3-amino-6-[3-(((biphenyl-4-ylamino)carbonyl)amino)methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
909	3-amino-6-(3-(((4-bromo-2-fluorophenyl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
910	N-[3-(5-amino-6-(((3S)-piperidin-3-ylamino)carbonyl)pyrazin-2-yl)phenyl]quinoline-3-carboxamide	A
911	3-amino-6-[3-(((6-(methyloxy)-1-benzofuran-3-yl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
912	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4'-(trifluoromethyl)biphenyl-2-yl)carbonyl)amino)phenyl]pyrazine-2-carboxamide	A
913	3-amino-6-[3-(((3,5-bis(trifluoromethyl)phenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
914	3-amino-6-[3-(((2,4-difluorophenyl)amino)carbonyl)amino)methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
915	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2-(trifluoromethyl)phenyl)sulfonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
916	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-((trifluoromethyl)oxy)phenyl)sulfonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
917	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2-((trifluoromethyl)oxy)phenyl)sulfonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
918	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(trifluoromethyl)phenyl)sulfonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B
919	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2-((trifluoromethyl)oxy)phenyl)amino)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide	B

#	Name	Chk1 IC ₅₀
920	3-amino-6-(3-(((2-(methylthio)phenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
921	3-amino-6-(3-(((3-bromo-5-methylphenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
922	3-amino-6-(3-(((2-bromophenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
923	5-[3-(aminomethyl)phenyl]-3-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine	A
924	3-amino-6-(3-(((4-(4-methylpiperazin-1-yl)phenyl)methyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
925	3-amino-6-(3-(((2-(4-fluorophenyl)ethyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
926	3-amino-6-(3-(((2-morpholin-4-ylethyl)amino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	B
927	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2R)-pyrrolidin-2-ylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide	A
928	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2S)-pyrrolidin-2-ylamino)carbonyl)phenyl)pyrazine-2-carboxamide	A
929	3-amino-6-(3-((cyclopentylamino)carbonyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	B
930	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-((trifluoromethyl)oxy)phenyl)amino)carbonyl)amino)methyl)phenyl]pyrazine-2-carboxamide	A
931	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((4-(trifluoromethyl)phenyl)amino)carbonyl)amino)methyl)phenyl]pyrazine-2-carboxamide	A
932	3-amino-6-(3-(((4-(methyloxy)phenyl)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
933	3-amino-6-(3-(((biphenyl-4-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
934	3-amino-6-(3-(((furan-2-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
935	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-ylamino)carbonyl)amino)methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
936	3-amino-6-(3-(((2,3-dihydro-1-benzofuran-5-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
937	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-6-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
938	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2-thienylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide	A
939	3-amino-6-(3-(((2,3-dihydro-1,4-benzodioxin-2-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
940	N-[[3-(5-amino-6-(3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl)pyrazin-2-yl)phenyl]methyl]-2-chlorobenzamide	A

#	Name	Chk1 IC ₅₀
941	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methyl]-4-chlorobenzamide	A
942	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methyl]-4-iodobenzamide	A
943	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methyl]-3,5-difluorobenzamide	A
944	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methyl]-4-bromo-2-fluorobenzamide	A
945	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methyl]-4-bromo-2-chlorobenzamide	A
946	3-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]-5-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenylpyrazin-2-amine	A
947	3-amino-N-azepan-3-yl-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenylpyrazine-2-carboxamide	A
948	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[2-(2-thienyl)-1,3-thiazol-4-yl]carbonyl]amino]methyl]phenylpyrazine-2-carboxamide	A
949	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[5-(2-thienyl)pyridin-3-yl]carbonyl]amino]methyl]phenylpyrazine-2-carboxamide	A
950	3-amino-6-[3-[[[4-(1,1-dimethylethyl)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
951	3-amino-6-[3-[[[2,4-dichloro-6-methylphenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
952	3-amino-6-[3-[[[3,5-dichlorophenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
953	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[2-thienylsulfonyl]amino]methyl)phenylpyrazine-2-carboxamide	A
954	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-[[3-thienylsulfonyl]amino]methyl)phenylpyrazine-2-carboxamide	A
955	3-amino-6-[3-[[[4-chlorophenyl]sulfonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
956	3-amino-6-[3-[[[1,3-benzodioxol-5-ylamino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
957	3-amino-6-(3-[[[4-(methyloxy)phenyl]amino]carbonyl]amino]methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
958	3-amino-6-[3-[[[4-fluorophenyl]amino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
959	3-amino-6-(3-[[2-(diethylamino)ethyl]oxy]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
960	3-amino-6-[3-[(2-morpholin-4-ylethyl)oxy]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	B
961	3-amino-6-[3-(methyloxy)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
962	3-amino-6-[3-(ethyloxy)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
963	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-(1,2,3-thiadiazol-4-yl)phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
964	3-amino-6-(3-[[2-(ethylamino)ethyl]oxy]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
965	phenylmethyl 4-(2-[[3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl]pyrazin-2-yl)phenyl]oxy)ethyl)piperazine-1-carboxylate	A
966	3-amino-6-(3-[[[4-(dimethylamino)phenyl]amino]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
967	3-amino-N-azepan-3-yl-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazine-2-carboxamide	A
968	3-amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
969	3-amino-6-[3-[[[(1S,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide	C
970	3-amino-N-[(3S)-azepan-3-yl]-6-[3-[[[(1S,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide	A
971	3-amino-6-[3-[[[4-(4-chlorophenyl)amino]carbonyl]amino]methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
972	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide	B
973	3-amino-N-[(3S)-azepan-3-yl]-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazine-2-carboxamide	A
974	3-amino-N-[(3S)-azepan-3-yl]-6-(3-methylphenyl)pyrazine-2-carboxamide	A
975	3-amino-6-(3-[[[2-(methylphenyl)acetyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
976	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[3-(trifluoromethyl)phenyl]acetyl]amino]phenyl]pyrazine-2-carboxamide	A
977	3-amino-6-[3-[[[4-(methyloxy)phenyl]acetyl]amino]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
978	3-amino-6-(3-[[[6-methylpyridin-3-yl]carbonyl]amino]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
979	3-amino-6-(3-[[[3-fluorophenyl]acetyl]amino]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
980	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[pyridin-3-ylacetyl]amino]phenyl]pyrazine-2-carboxamide	A
981	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[pyridin-3-ylcarbonyl]amino]phenyl]pyrazine-2-carboxamide	A
982	3-amino-6-(3-[[[2,5-difluorophenyl]acetyl]amino]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
983	3-amino-6-(3-[[[4-fluorophenyl]acetyl]amino]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
984	3-amino-6-[3-[[[3-fluoro-5-(trifluoromethyl)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
985	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
986	3-Amino-6-[3-(((5-(4-nitro-phenyl)-furan-2-carbonyl)-amino)-methyl)-phenyl]-pyrazine-2-carboxylic acid (3S)-piperidin-3-ylamide	A
987	3-amino-6-[3-(((3-hydroxypyridin-2-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
988	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(trifluoromethyl)oxy)phenyl)oxy)acetyl]amino)methyl]phenyl]pyrazine-2-carboxamide	A
989	3-amino-6-[3-(((5-methylisoxazol-3-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
990	3-amino-6-[3-(((isoxazol-5-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
991	3-amino-6-[3-(((4-methyl-1,2,3-thiadiazol-5-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
992	3-amino-6-[3-(((2,5-dichloro-3-thienyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
993	3-amino-6-[3-(((5-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
994	3-amino-6-[3-(((2-chloro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
995	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
996	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
997	3-amino-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
998	3-amino-6-[3-(((2-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
999	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1000	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1001	3-amino-6-[3-(((2-fluoro-4-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1002	3-amino-6-[3-(((2-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1003	3-amino-6-[3-(((2,3-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1004	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1005	3-amino-6-[3-(((4-chloro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1006	3-amino-6-[3-(((3-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
1007	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1008	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1009	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1010	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1011	3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1012	3-amino-6-[3-(((6-chloro-2-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1013	3-amino-6-[3-(((2-chloro-6-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1014	3-amino-6-[3-(((2-chloro-3,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1015	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1016	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,4-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1017	3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1018	3-amino-6-[3-(((3-fluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1019	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1020	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1021	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1022	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1023	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1024	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1025	3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1026	3-amino-6-[3-(((2,6-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	B
1027	3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1028	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A

#	Name	Chk1 IC ₅₀
1029	3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1030	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1031	3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1032	3-amino-6-[3-(((3-bromo-5-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1033	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide	A
1034	3-amino-6-[3-(((biphenyl-3-yl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1035	3-amino-6-[3-(((2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1036	3-amino-6-[3-(((2,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1037	3-amino-6-[3-(((3,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1038	3-amino-6-[3-(((2,4-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1039	3-amino-6-[3-(((3-methoxyphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1040	3-amino-6-[3-(((phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1041	3-amino-6-[3-(((3,5-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A
1042	3-amino-6-[3-(((3,4-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A

[1448] Table 12 shows structure activity relationship data for selected compounds of the invention with respect to Chk1 and a number of other enzymes, thus indicating selectivity of exemplary compounds of the invention toward Checkpoint kinases, particularly Chk1. Inhibition is indicated as IC₅₀ with following key: A = IC₅₀ less than 50 nM, B = IC₅₀ greater than 50 nM but less than or equal to 1000 nM, C = IC₅₀ greater than 1000 nM but less than 10,000 nM, and D = IC₅₀ of 10,000 nM or greater (a blank cell in the table indicates lack of data only).

[1449] Abbreviations for human enzymes listed in Table 12 are defined as follows: EphB4 refers to ephrin-related receptor tyrosine kinase B4; Tie-2 refers to tyrosine kinase with immunoglobulin and EGF repeats; EGFR refers to epidermal growth factor receptor; FGFR1 refers to fibroblast growth factor receptor; KDR refers to kinase insert domain

containing receptor; Flt1 refers to fms-like tyrosine kinase 1; fyn refers to proto-oncogene tyrosine-protein kinase fyn; and PDGFR- β refers to platelet derived growth factor receptor beta chain. Chk1 refers to checkpoint homologue 1 (*S. pombe*); Chk2 refers to checkpoint-like protein 2; Flt1 refers to fms-like tyrosine kinase 1, Flt4 refers to fms-like tyrosine kinase 4; fyn refers to proto-oncogene tyrosine protein kinase fyn; KDR refers to kinase insert domain containing receptor; MAP4K3 refers to mitogen-activated protein kinase kinase kinase kinase 3; c-MET refers to the cellular homologue of the met proto-oncogene; CLK1 refers to CDC-like kinase1; EMK refers to ELKL motif kinase 1; ErbB2 refers to erythroblastic leukemia viral Oncogene homologue 2; IGFR1 refers to insulin-like growth factor receptor 1; IRK refers to insulin receptor kinase; MAPKAP2 refers to mitogen-activated protein kinase-activated protein kinase; PKA refers to protein kinase A; PKC refers to protein kinase C epsilon isoform; PLK refers to polo-like kinase; scr refers to the src proto-oncogene; ZAP70 refers to zeta-chain associated protein kinase, 70kD; cdc2 refers to cell division cycle protein 2.

Table 12

#	Compound	Chk1	Chk2	FGFR1	Flt1	Flt4	Fyn	PDGFR- β	KDR	MAP4K3	c-Met	CLK1	EGFR	EMK	EphA2R	EphB4	ErbB2	IGFR1	IRK	MAPKAP2	PKA	PKC ϵ	PLK	SRC	Tie-2	ZAP70	Cdc2
1	3-amino-6-[3-(((4-(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	C	B	B	C	B	B	B		C	D	D	D	C	D		D		C	D	D				D
2	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((4-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide	A	A	C	B	A	C	B	B	A		C	D	C	D	D	D		D		C	D	C				D
3	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	C	B	A	B	B	B	A		C	D	C	D	C	D		D		C	D	D				D

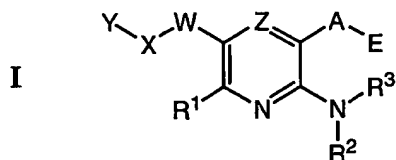
#	Compound	Chk1	Chk2	FGFR1	Flt1	Flt4	Fyn	PDGFR-β	KOR	MAP4K3	c-Met	CLK1	EGFR	EMK	EphA2R	EphB4	ErbB2	IGFR1	IRK	MAPKAP2	PKA	PKCe	PLK	SRC	Tle-2	ZAP70	Cdc2
4	3-amino-6-[3-(((4-(dimethylamino)phenyl)methyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	C	B	B	B	B	B	B		C	D	D	D	D	C		D		D	D	D				D
5	3-amino-6-[3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	C	B	B	B	B	B	A		B	D	C	D	C	D		D		C	D	D				D
6	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	C	B	A	C	A	B	B	C	C		D	C	C	D	D	D	C	D	D	D	C	C	C	D
7	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	C	B	A	C	B	A	A		C	D	C	D	C	D		D		C	D	D				D
8	3-amino-6-[3-(((4-fluorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	B	B	A	B	B	B	A		D	C	C	C	C	D		D		C	D	D				
9	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	B	B	A	B	A	A	A	C	C		C	C	C	D	D	D	C	C	C	D	B	B	C	C
10	3-amino-6-[3-(((4-chlorophenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	B	A	A	B	A	A	A	C	C		C	C	C	D	D	D	C	C	C	D	B	B	B	C

#	Compound	Chk1	Chk2	FGFR1	Flt1	Flt4	Fyn	PDGFR-β	KDR	MAPK3	c-Met	CLK1	EGFR	EMK	EphA2R	EphB4	ErbB2	IGFR1	IRK	MAPKAP2	PKA	PKCε	PLK	SRC	Tie-2	ZAP70	Cdc2
11	3-amino-6-(3- {[(pentafluorophenyl)acet yl]amino}phenyl)-N- [(3S)-piperidin-3- yl]pyrazine-2- carboxamide	A	A	B	B	A	B	B	A	A		C	C	C	D	C	D		D		C	D	D				
12	3-amino-6-(3- {[(2,4- difluorophenyl)acetyl]ami no}phenyl)-N-[(3S)- piperidin-3-yl]pyrazine-2- carboxamide	A	A	B	B	A	B	A	A	A	C	C	B	C	C	C	C	D	D	C	C	C	D	B	B	C	D
13	3-amino-6-[3- {[(biphenyl-4- ylamino)carbonyl]amino} methyl}phenyl]-N-[(3S)- piperidin-3-yl]pyrazine-2- carboxamide	A	A	B	B	A	B	A	A	A		D	D	C	D	C	C		D		B	D	C				
14	3-amino-6-(3- {[(4-bromo- 2- fluorophenyl)carbonyl]am ino}phenyl)-N-[(3S)- piperidin-3-yl]pyrazine-2- carboxamide	A	A	B	A	A	B	A	A	A	C	C		C	C	C	D	D	D	C	C	C	C	C	C	B	D
15	N-[3-(5-amino-6- {[(3S)- piperidin-3- ylamino]carbonyl}pyrazin -2-yl)phenyl]quinoline-3- carboxamide	A	A	B	A	A	B	A	A	A		B	C	C	D	C	D		D		C	D	D				
16	3-amino-6-(3- {[(biphenyl- 4- ylcarbonyl)amino]methyl} phenyl)-N-[(3S)-piperidin- 3-yl]pyrazine-2- carboxamide	A	A	B	B	A	B	A	A	A	C	C		C	C	B	D	D	D	C	B	D	C	B	C	C	C
17	3-amino-N-[(3S)- piperidin-3-yl]-6-(3- {[(2- thienylcarbonyl)amino]me thyl}phenyl)pyrazine-2- carboxamide	A	A	B	B	A	B	A	A	A		D	C	C	D	C	D		D		C	D	D				
18	3-amino-6-(3- {[2- (ethylamino)ethyl]oxy}ph enyl)-N-[(3S)-piperidin-3- yl]pyrazine-2- carboxamide	A	A	B	B	A	B	B	B	B		C	C	C	C	C	D		D		D	D	D				

#	Compound	Chk1	Chk2	FGFR1	Flt1	Flt4	Fyn	PDGFR- β	KDR	MAP4K3	c-Met	CLK1	EGFR	EMK	EphA2R	EphB4	ErbB2	IGFR1	IRK	MAPKAP2	PKA	PKC α	PLK	SRC	Tle-2	ZAP70	Cdc2
19	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide	A	A	B	B	A	B	A	A	A		D	C	B	D	C	D		D		C	D	D				

What is claimed is:

1. A compound for modulating protein kinase activity of Formula I,



or a pharmaceutically acceptable salt, hydrate, or prodrug thereof, wherein,

R^1 is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl;

R^2 and R^3 are each independently selected from -H, optionally substituted alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl; or

R^2 and R^3 , together with the nitrogen to which they are attached, are combined to form an optionally substituted five- to seven-membered heterocyclyl, said five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

R^4 is -H or R^5 ;

R^5 is selected from optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl; or

R^4 and R^5 , when taken together with a common nitrogen to which they are attached, form an optionally substituted five- to seven-membered heterocyclyl, said optionally substituted five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

Z is -N= or -C(H)=;

A is selected from -C(=O)-, -C(=S)-, -C(=NR⁶)-, and -R⁷, when A is -R⁷, E does not exist;

R^6 is selected from -H, -NO₂, -N(R²)R³, -NC(=NR²)N(R²)R³, -CN, -OR⁴, optionally substituted lower alkyl, optionally substituted heteroalicycylalkyl, optionally substituted aryl, optionally substituted arylalkyl and optionally substituted heteroalicyclic;

R^7 is an optionally substituted five- to seven-membered heterocyclyl, said five- to seven-membered heterocyclyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

E is selected from -N(R⁸)R⁹-, -NN(R²)R³-, -NNC(=O)N(R²)R³-, -OR⁴, and -R¹⁰;

R^8 is -H or optionally substituted alkyl;

R^9 is selected from -H, optionally substituted heteroalicycylalkyl, optionally substituted heteroarylalkyl, and optionally substituted heteroalicyclic; or

R^8 and R^9 , together with the nitrogen to which they are attached, are combined to form an optionally substituted five- to seven-membered heteroalicycyl, said five- to seven-membered heteroalicycyl optionally containing at least one additional heteroatom selected from N, O, S, and P;

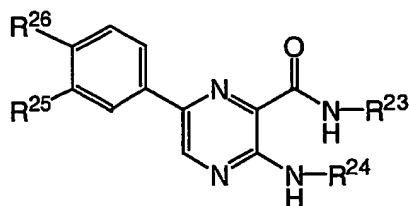
R^{10} is selected from -H, optionally substituted heteroalicycylalkyl, optionally substituted heteroalicyclic, optionally substituted arylalkyl, optionally substituted aryl, and optionally substituted alkyl;

W is either an optionally substituted six- to ten-membered arylene or an optionally substituted five- to ten-membered heteroarylene;

X is selected from a single bond, -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₀₋₃-, -(CH₂)₀₋₃CO₂(CH₂)₀₋₃-, -(CH₂)₀₋₃SO₂N(R⁴)(CH₂)₀₋₃-, -N(R⁴)(CH₂)₂₋₃O-, -(CH₂)₀₋₃N(R⁴)C(=O)N(R⁴)(CH₂)₀₋₃-, -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₂₋₃O-, -(CH₂)₀₋₃S(O)₀₋₂(CH₂)₀₋₃-, -N(R⁴)(CH₂)₂₋₃N(R⁴)-, -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₁₋₃C(=O)-, -(CH₂)₀₋₃OC(=O)N(R⁴)(CH₂)₀₋₃-, -C(=O)N(R⁴)N(R⁴)-, -(CH₂)₀₋₃CO₂(CH₂)₂₋₃N(R⁴)-, -(CH₂)₀₋₃N(R⁴)C(=O)(CH₂)₀₋₃O-, -(CH₂)₀₋₃N(R⁴)(CH₂)₀₋₃-, -C(=O)N(R⁴)(CH₂)₂₋₃N(R⁴)-, -O(CH₂)₂₋₃O-, -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₂₋₃S(O)₀₋₂-, -(CH₂)₀₋₃N(R⁴)C(=O)(CH₂)₀₋₃S(O)₀₋₂-, -(CH₂)₀₋₃O(CH₂)₀₋₃-, optionally substituted alkoxy, and optionally substituted lower alkylene, wherein any -CH₂- in X is optionally substituted; and

Y is selected from -H, optionally substituted lower alkyl, optionally substituted aryl, and optionally substituted heterocyclyl;

provided the compound is not:



wherein R^{23} , R^{24} , and R^{25} are -H; and R^{26} is -H, -Cl, -CH₃, or -OCH₃; or

wherein R^{23} is -C(NH)NH₂; R^{24} is -CH₃; R^{25} is H; and R^{26} is -Cl; or

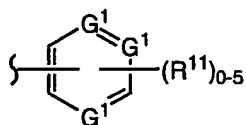
wherein R^{23} is -H, methyl, ethyl, propyl, isopropyl, butyl, t-butyl, 1-methylpropyl, hydroxyethyl; and R^{24} , R^{25} , and R^{26} are H; or

wherein R^{23} is -H; R^{24} is butyl; and R^{25} and R^{26} are methoxy; or

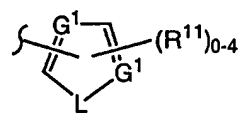
wherein R^{23} 1-methylpropyl; R^{24} is -CHO; R^{25} is -H; and R^{26} is -Cl; or

wherein R^{23} is -H; R^{24} is -CH₃; and R^{25} and R^{26} are -H.

2. The compound according to claim 1, wherein Z is =N-.
3. The compound according to claim 2, wherein R^1 is -H.
4. The compound according to claim 3, wherein R^2 is -H and R^3 is either -H or lower alkyl.
5. The compound according to claim 4, wherein R^2 and R^3 are -H.
6. The compound according to claim 5, wherein A is R^7 , and R^7 is either

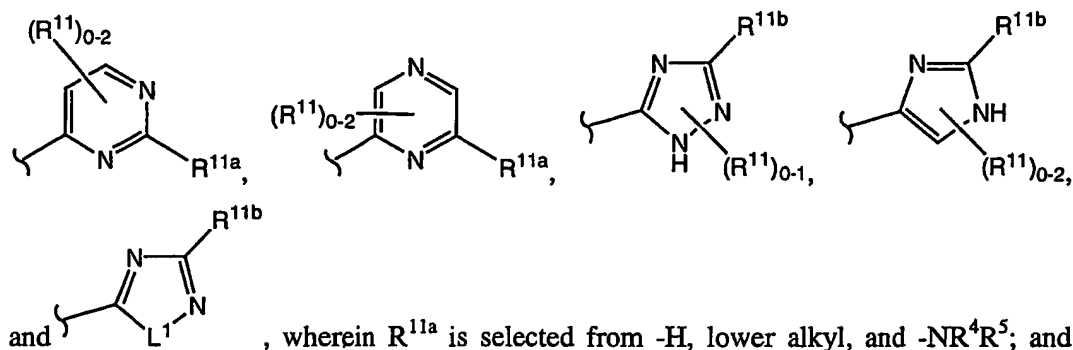


or



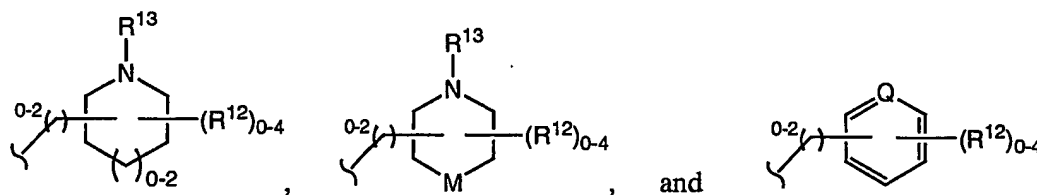
wherein each G^1 is independently =N- or =C(R^{11})-, L is selected from -O-, -S(O)₀₋₂-, and -NR¹¹, wherein each R^{11} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R^4)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R^4)R⁵, -CO₂R⁴, -C(O)N(R^4)R⁵, -N(R^4)SO₂R⁵, -N(R^4)C(O)R⁵, -N(R^4)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; or optionally two R^{11} 's form a ring system fused with the existing ring of R^7 , said ring system substituted with 0 to 3 additional of R^{11} and said ring system optionally containing between 1 and 3 heteroatoms.

7. The compound according to claim 6, wherein R^7 is selected from



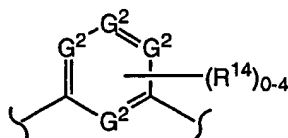
and $(R^{11})_{0-2}$, wherein R^{11a} is selected from -H, lower alkyl, and $-NR^4R^5$; and R^{11b} is selected from optionally substituted alkyl, optionally substituted aryl, optionally substituted arylalkyl, optionally substituted heteroaryl, optionally substituted heteroarylalkyl, and optionally substituted heteroalicyclic; and L is selected from -O-, $-S(O)_{0-2}$.

8. The compound according to claim 7, wherein R^{11b} is selected from optionally substituted alkyl,



wherein each R^{12} is independently selected from -H, halogen, oxo, -CN, $-NH_2$, $-CF_3$, $-NO_2$, $-OR^4$, $-N(R^4)R^5$, $-S(O)_{0-2}R^5$, $-SO_2N(R^4)R^5$, $-CO_2R^4$, $-C(O)N(R^4)R^5$, $-N(R^4)SO_2R^5$, $-N(R^4)C(O)R^5$, $-N(R^4)CO_2R^5$, $-C(O)R^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; R^{13} is selected from -H, lower alkyl, and acyl; Q is either $=N-$ or $=C(R^{12})-$; and M is selected from -O-, $-S(O)_{0-2}$, and $-NR^{13}$.

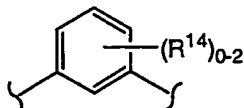
9. The compound according to claim 8, wherein W is



wherein each G^2 is independently $=N-$ or $=C(R^{14})-$, wherein each R^{14} is independently selected from -H, halogen, -CN, $-NH_2$, $-CF_3$, $-NO_2$, $-OR^4$, $-N(R^4)R^5$, $-S(O)_{0-2}R^5$, $-SO_2N(R^4)R^5$, $-CO_2R^4$, $-C(O)N(R^4)R^5$, $-N(R^4)SO_2R^5$, $-N(R^4)C(O)R^5$, $-N(R^4)CO_2R^5$, $-C(O)R^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally

substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

10. The compound according to claim 9, wherein W is



11. The compound according to claim 10, wherein X is selected from $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}SO_2N(R^4)(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}N(R^4)C(=O)N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)N(R^4)-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}O-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{1-3}C(=O)-$, $-O(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}OC(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}O(CH_2)_{0-3}-$, $-(CH_2)_{0-3}CO_2(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}O-$, $-C(=O)N(R^4)(CH_2)_{2-3}N(R^4)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

12. The compound according to claim 11, wherein X is selected from $-C(=O)N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}C(=O)N(H)-$, $-C(=O)N(H)-$, $-SO_2N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}SO_2N(H)-$, $-SO_2N(H)-$, $-N(H)C(=O)N(H)(CH_2)_{1-2}-$, $-N(H)C(=O)N(H)-$, $-(CH_2)_{0-2}N(R^4)(CH_2)_{0-2}-$, $-(CH_2)_{0-2}O(CH_2)_{0-2}-$, $-C(=O)N(H)N(H)-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-OCH_2C(=O)N(H)CH_2-$, $-(CH_2)_{1-2}N(H)C(=O)N(H)-$, $-OCH_2CH_2O-$, $-N(R^4)CH_2C(=O)N(H)CH_2-$, $-(CH_2)_{0-1}C(=O)N(H)(CH_2)_{1-2}C(=O)-$, optionally substituted alkoxy, and optionally substituted lower alkylene.

13. The compound according to claim 12, wherein Y is selected from optionally substituted cycloalkyl, optionally substituted phenyl, and optionally substituted five- to six-membered heterocyclyl.

14. The compound according to claim 13, wherein Y is selected from Table 1.

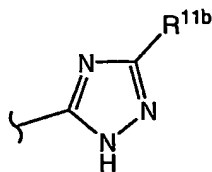
Table 1

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wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl.

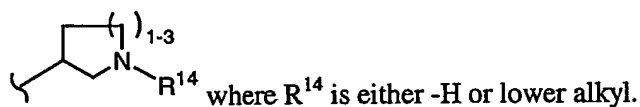
15. The compound according to claim 14, wherein R^{11b} is selected from optionally substituted alkyl, optionally substituted phenyl, optionally substituted pyridyl, and optionally substituted heteroalicyclic.

16. The compound according to claim 15, wherein R^7 is



17. The compound according to claim 16, wherein R^{11b} is selected from optionally substituted morpholine, optionally substituted pyrrolidine, optionally substituted piperidine, and optionally substituted azepane.

18. The compound according to claim 17, wherein R^{11b} is



19. The compound according to claim 16, wherein R^{11b} is a pyridine.

20. The compound according to claim 19, wherein R^{11b} is a pyridine bearing a 2-amino group.

21. The compound according to claim 4, wherein A is $-C(=O)-$ or $-C(=NR^6)-$.

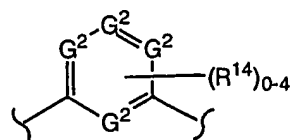
22. The compound according to claim 21, wherein A is $-C(=O)-$.

23. The compound according to claim 22, wherein E is selected from Table 2.

Table 2

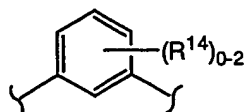
wherein each R^{16} is independently selected from -H, optionally substituted lower alkyl, $-\text{CO}_2R^4$, and $-\text{C}(=\text{O})R^4$; or optionally two of R^{16} , together with the nitrogen or nitrogens to which they are attached, form a heterocyclic ring; each R^{17} is independently selected from -H, halogen, oxo, -CN, $-\text{NH}_2$, $-\text{CF}_3$, $-\text{NO}_2$, $-\text{OR}^4$, $-\text{N}(R^4)R^5$, $-\text{S}(\text{O})_{0-2}R^5$, $-\text{SO}_2\text{N}(R^4)R^5$, $-\text{CO}_2R^4$, $-\text{C}(\text{O})\text{N}(R^4)R^5$, $-\text{N}(R^4)\text{SO}_2R^5$, $-\text{N}(R^4)\text{C}(\text{O})R^5$, $-\text{N}(R^4)\text{CO}_2R^5$, $-\text{C}(\text{O})R^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl; R^{18} is optionally substituted lower alkyl; J^1 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0]; and J^2 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0], wherein the nitrogen bearing R^{16} and any nitrogen in J^2 must have at least two carbons between them.

24. The compound according to claim 23, wherein W is



wherein each G^2 is independently =N- or =C(R¹⁴)-, wherein each R¹⁴ is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclylalkyl.

25. The compound according to claim 24, wherein W is



26. The compound according to claim 25, wherein X is selected from -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₀₋₃-, -(CH₂)₀₋₃SO₂N(R⁴)(CH₂)₀₋₃-, -N(R⁴)(CH₂)₂₋₃O-, -(CH₂)₀₋₃N(R⁴)C(=O)N(R⁴)(CH₂)₀₋₃-, -C(=O)N(R⁴)N(R⁴)-, -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₂₋₃O-, -N(R⁴)(CH₂)₂₋₃N(R⁴)-, -(CH₂)₀₋₃N(R⁴)(CH₂)₀₋₃-, -(CH₂)₀₋₃C(=O)N(R⁴)(CH₂)₁₋₃C(=O)-, -O(CH₂)₂₋₃O-, -(CH₂)₀₋₃OC(=O)N(R⁴)(CH₂)₀₋₃-, -(CH₂)₀₋₃O(CH₂)₀₋₃-, -(CH₂)₀₋₃CO₂(CH₂)₂₋₃N(R⁴)-, -(CH₂)₀₋₃N(R⁴)C(=O)(CH₂)₀₋₃O-, -C(=O)N(R⁴)(CH₂)₂₋₃N(R⁴)-, optionally substituted alkoxyl, and optionally substituted lower alkylene.

27. The compound according to claim 26, wherein X is selected from -C(=O)N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂C(=O)N(H)-, -C(=O)N(H)-, -SO₂N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂SO₂N(H)-, -SO₂N(H)-, -N(H)C(=O)N(H)(CH₂)₁₋₂-, -N(H)C(=O)N(H)-, -(CH₂)₀₋₂N(R⁴)(CH₂)₀₋₂-, -(CH₂)₀₋₂O(CH₂)₀₋₂-, -C(=O)N(H)N(H)-, -N(R⁴)(CH₂)₂₋₃N(R⁴)-, -OCH₂C(=O)N(H)CH₂-, -(CH₂)₁₋₂N(H)C(=O)N(H)-, -OCH₂CH₂O-, -N(R⁴)CH₂C(=O)N(H)CH₂-, -(CH₂)₀₋₁C(=O)N(H)(CH₂)₁₋₂C(=O)-, optionally substituted alkoxyl, and optionally substituted lower alkylene.

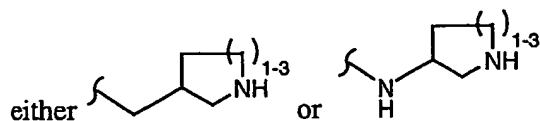
28. The compound according to claim 27, wherein Y is selected from optionally substituted cycloalkyl, optionally substituted phenyl, and optionally substituted five- to six-membered heterocyclyl.

29. The compound according to claim 28, wherein Y is selected from Table 3.

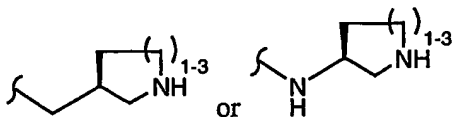
Table 3

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

30. The compound according to claim 29, wherein E is



31. The compound according to claim 30, wherein E is either

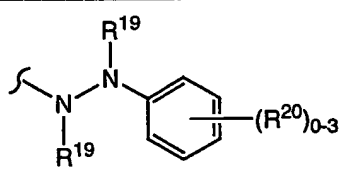
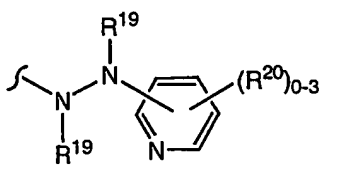
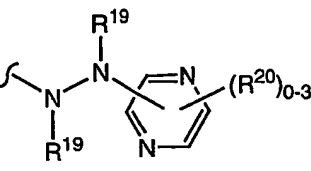
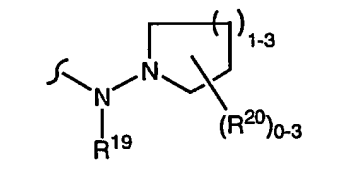
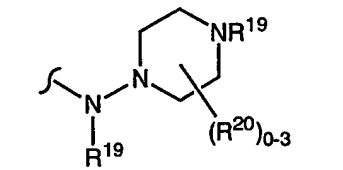
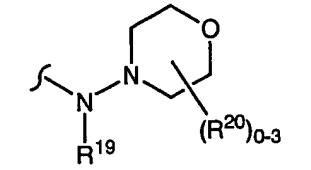
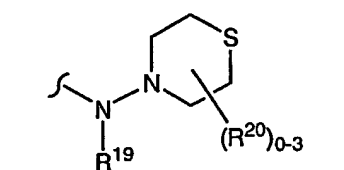
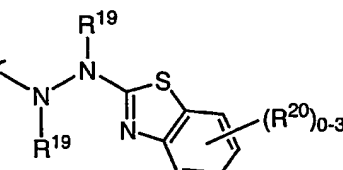
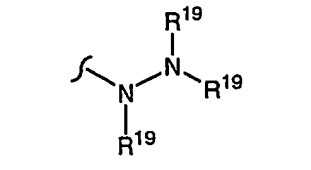
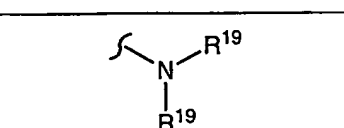


32. The compound according to claim 31, wherein R^2 and R^3 are -H.

33. The compound according to claim 21, wherein A is $-C(=NR^6)-$.

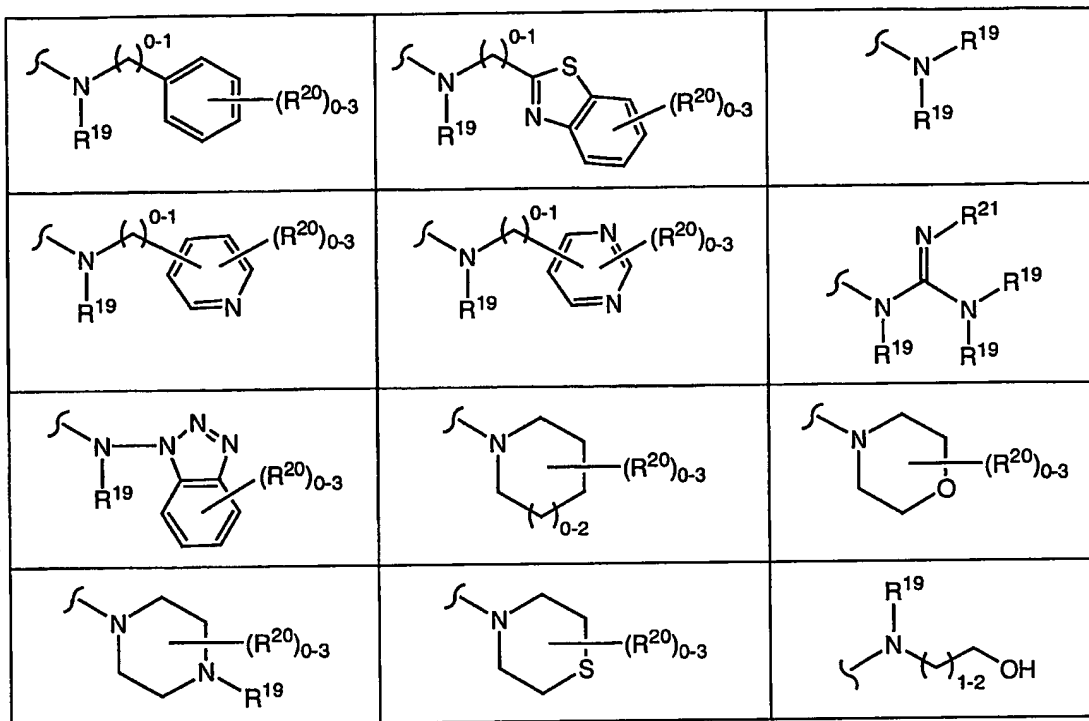
34. The compound according to claim 33, wherein E is selected from -H, lower alkyl, and a group listed in Table 4,

Table 4

R^6 is selected from -H, -OH, lower alkyl, -NO₂, -CN, lower alkoxy, and a group listed in Table 5,

Table 5

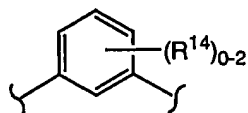


wherein each R^{19} is independently either $-H$ or lower alkyl;

each R^{20} is independently selected from $-H$, halogen, oxo, $-CN$, $-NH_2$, $-CF_3$, $-NO_2$, $-OR^4$, $-N(R^4)R^5$, $-S(O)_{0-2}R^5$, $-SO_2N(R^4)R^5$, $-CO_2R^4$, $-C(O)N(R^4)R^5$, $-N(R^4)SO_2R^5$, $-N(R^4)C(O)R^5$, $-N(R^4)CO_2R^5$, $-C(O)R^4$, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl;

and R^{21} is selected from $-H$, $-OH$, lower alkyl, $-NO_2$, $-CN$, lower alkoxy.

35. The compound according to claim 34, wherein W is



36. The compound according to claim 35, wherein X is selected from $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}SO_2N(R^4)(CH_2)_{0-3}-$, $-N(R^4)(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}N(R^4)C(=O)N(R^4)(CH_2)_{0-3}-$, $-C(=O)N(R^4)N(R^4)-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{2-3}O-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}C(=O)N(R^4)(CH_2)_{1-3}C(=O)-$, $-O(CH_2)_{2-3}O-$, $-(CH_2)_{0-3}OC(=O)N(R^4)(CH_2)_{0-3}-$, $-(CH_2)_{0-3}O(CH_2)_{0-3}-$, $-(CH_2)_{0-3}CO_2(CH_2)_{2-3}N(R^4)-$, $-(CH_2)_{0-3}N(R^4)C(=O)(CH_2)_{0-3}O-$,

-C(=O)N(R⁴)(CH₂)₂₋₃N(R⁴)-, optionally substituted alkoxy, and optionally substituted lower alkylene.

37. The compound according to claim 36, wherein X is selected from -C(=O)N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂C(=O)N(H)-, -C(=O)N(H)-, -SO₂N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂SO₂N(H)-, -SO₂N(H)-, -N(H)C(=O)N(H)(CH₂)₁₋₂-, -N(H)C(=O)N(H)-, -(CH₂)₀₋₂N(R⁴)(CH₂)₀₋₂-, -(CH₂)₀₋₂O(CH₂)₀₋₂-, -C(=O)N(H)N(H)-, -N(R⁴)(CH₂)₂₋₃N(R⁴)-, -OCH₂C(=O)N(H)CH₂-, -(CH₂)₁₋₂N(H)C(=O)N(H)-, -OCH₂CH₂O-, -N(R⁴)CH₂C(=O)N(H)CH₂-, -(CH₂)₀₋₁C(=O)N(H)(CH₂)₁₋₂C(=O)-, optionally substituted alkoxy, and optionally substituted lower alkylene.

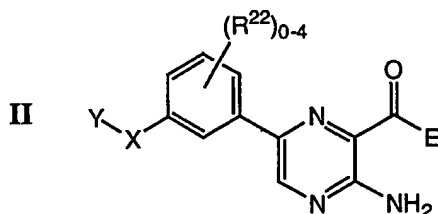
38. The compound according to claim 37, wherein Y is selected from optionally substituted cycloalkyl, optionally substituted phenyl, and optionally substituted five- to six-membered heterocyclyl.

39. The compound according to claim 38, wherein Y is selected from Table 6.

Table 6

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

40. The compound according to claim 1, of formula II,



wherein E is selected from Table 7,

Table 7

wherein each R^{16} is independently selected from -H, optionally substituted lower alkyl, $-CO_2R^4$, and $-C(=O)R^4$;

or optionally two of R^{16} , together with the nitrogen or nitrogens to which they are attached, form a heterocyclic ring;

each R^{17} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl;

R^{18} is optionally substituted lower alkyl;

J^1 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of [4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0];

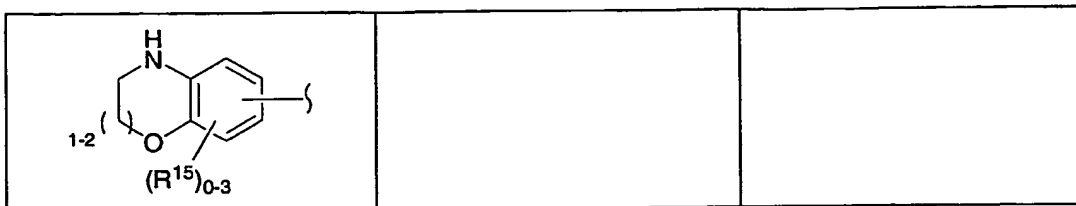
J^2 is a saturated bridged ring system containing at least one nitrogen ring atom, said saturated bridged ring system having a geometry selected from the group consisting of

[4.4.0], [4.3.0], [4.2.0], [4.1.0], [3.3.0], [3.2.0], and [3.1.0], wherein the nitrogen bearing R^{16} and any nitrogen in J^2 must have at least two carbons between them.

X is selected from $-C(=O)N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}C(=O)N(H)-$, $-C(=O)N(H)-$, $-SO_2N(H)(CH_2)_{1-2}-$, $-(CH_2)_{1-2}SO_2N(H)-$, $-SO_2N(H)-$, $-N(H)C(=O)N(H)(CH_2)_{1-2}-$, $-N(H)C(=O)N(H)-$, $-(CH_2)_{0-2}N(R^4)(CH_2)_{0-2}-$, $-(CH_2)_{0-2}O(CH_2)_{0-2}-$, $-C(=O)N(H)N(H)-$, $-N(R^4)(CH_2)_{2-3}N(R^4)-$, $-OCH_2C(=O)N(H)CH_2-$, $-(CH_2)_{1-2}N(H)C(=O)N(H)-$, $-OCH_2CH_2O-$, $-N(R^4)CH_2C(=O)N(H)CH_2-$, $-(CH_2)_{0-1}C(=O)N(H)(CH_2)_{1-2}C(=O)-$, optionally substituted alkoxy, and optionally substituted lower alkylene;

Y is selected from Table 8,

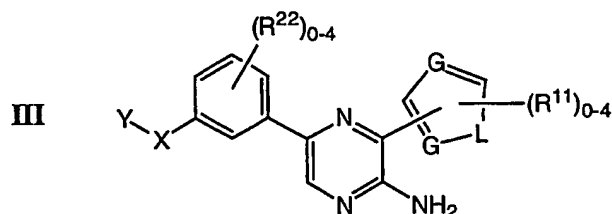
Table 8



wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl; and,

each R^{22} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

41. The compound according to claim 1, of formula **III**,



wherein, each G is independently =N- or =C(R¹¹)-;

L is selected from -O-, -S(O)₀₋₂-, and -NR¹¹;

each R^{11} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl; or

optionally two R^{11} 's form a ring system fused with the existing ring of R^7 , said ring system substituted with 0 to 3 additional of R^{11} and said ring system optionally containing between 1 and 3 heteroatoms;

X is selected from -C(=O)N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂C(=O)N(H)-, -C(=O)N(H)-, -SO₂N(H)(CH₂)₁₋₂-, -(CH₂)₁₋₂SO₂N(H)-, -SO₂N(H)-, -N(H)C(=O)N(H)(CH₂)₁₋₂-,

-N(H)C(=O)N(H)-, $-(\text{CH}_2)_{0-2}\text{N}(\text{R}^4)(\text{CH}_2)_{0-2}-$, $-(\text{CH}_2)_{0-2}\text{O}(\text{CH}_2)_{0-2}-$, $-\text{C}(=\text{O})\text{N}(\text{H})\text{N}(\text{H})-$,
 $-\text{N}(\text{R}^4)(\text{CH}_2)_{2-3}\text{N}(\text{R}^4)-$, $-\text{OCH}_2\text{C}(=\text{O})\text{N}(\text{H})\text{CH}_2-$, $-(\text{CH}_2)_{1-2}\text{N}(\text{H})\text{C}(=\text{O})\text{N}(\text{H})-$, $-\text{OCH}_2\text{CH}_2\text{O}-$,
 $-\text{N}(\text{R}^4)\text{CH}_2\text{C}(=\text{O})\text{N}(\text{H})\text{CH}_2-$, $-(\text{CH}_2)_{0-1}\text{C}(=\text{O})\text{N}(\text{H})(\text{CH}_2)_{1-2}\text{C}(=\text{O})-$, optionally
 substituted alkoxy, and optionally substituted lower alkylene;

Y is selected from Table 9,

Table 9

wherein each R^{15} is independently selected from -H, halogen, oxo, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl; and,

each R^{22} is independently selected from -H, halogen, -CN, -NH₂, -CF₃, -NO₂, -OR⁴, -N(R⁴)R⁵, -S(O)₀₋₂R⁵, -SO₂N(R⁴)R⁵, -CO₂R⁴, -C(O)N(R⁴)R⁵, -N(R⁴)SO₂R⁵, -N(R⁴)C(O)R⁵, -N(R⁴)CO₂R⁵, -C(O)R⁴, optionally substituted lower alkyl, optionally substituted aryl, optionally substituted lower arylalkyl, optionally substituted heterocyclyl, and optionally substituted lower heterocyclalkyl.

42. The compound according to claim 1, wherein the compound is selected from Table 10.

Table 10

#	Name
1	3-amino-6-phenyl-N-(2-phenylethyl)pyrazine-2-carboxamide
2	3-amino-6-phenyl-N-(phenylmethyl)pyrazine-2-carboxamide
3	3-amino-6-(3-chlorophenyl)pyrazine-2-carboxamide
4	3-amino-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide
5	3-amino-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-6-phenylpyrazine-2-carboxamide
6	3-amino-N-(2,3-dihydro-1H-inden-2-yl)-6-phenylpyrazine-2-carboxamide
7	3-amino-6-(2-chlorophenyl)pyrazine-2-carboxamide
8	3-amino-N-cyclopentyl-6-phenylpyrazine-2-carboxamide
9	3-amino-N,N-dimethyl-6-phenylpyrazine-2-carboxamide
10	3-amino-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide
11	5-phenyl-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine
12	3-amino-N-(cyclopropylmethyl)-6-phenylpyrazine-2-carboxamide
13	3-amino-6-phenyl-N-(tetrahydrofuran-2-ylmethyl)pyrazine-2-carboxamide

#	Name
14	3-amino-N-cyclohexyl-6-phenylpyrazine-2-carboxamide
15	6-phenyl-N-propyl-3-(propylamino)pyrazine-2-carboxamide
16	3-amino-6-(2-methylphenyl)pyrazine-2-carboxamide
17	3-amino-6-[2-(methyloxy)phenyl]pyrazine-2-carboxamide
18	3-amino-6-(2,4-difluorophenyl)pyrazine-2-carboxamide
19	3-amino-6-(3-fluorophenyl)pyrazine-2-carboxamide
20	3-amino-N-cyclopropyl-6-phenylpyrazine-2-carboxamide
21	3-amino-6-(3,4-difluorophenyl)pyrazine-2-carboxamide
22	3-amino-6-(4-fluorophenyl)-N-methylpyrazine-2-carboxamide
23	3-amino-6-[4-(ethyloxy)phenyl]pyrazine-2-carboxamide
24	3-amino-6-[3-(ethyloxy)phenyl]pyrazine-2-carboxamide
25	3-amino-6-(1,3-benzodioxol-5-yl)pyrazine-2-carboxamide
26	3-amino-6-naphthalen-1-ylpyrazine-2-carboxamide
27	3-amino-6-naphthalen-2-ylpyrazine-2-carboxamide
28	3-amino-6-{3-[(trifluoromethyl)oxy]phenyl}pyrazine-2-carboxamide
29	3-amino-6-[3-(aminocarbonyl)phenyl]pyrazine-2-carboxamide
30	6-[3-(acetylamino)phenyl]-3-aminopyrazine-2-carboxamide
31	3-amino-6-biphenyl-4-ylpyrazine-2-carboxamide
32	3-amino-6-[4-(dimethylamino)phenyl]pyrazine-2-carboxamide
33	3-amino-6-(3-methylphenyl)pyrazine-2-carboxamide
34	methyl 3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoate
35	3-amino-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide
36	3-amino-6-(3-hydroxyphenyl)pyrazine-2-carboxamide

#	Name
37	3-amino-6-(1-benzofuran-2-yl)pyrazine-2-carboxamide
38	3-(methylamino)-6-phenylpyrazine-2-carboxamide
39	6-phenyl-3-[(phenylmethyl)amino]pyrazine-2-carboxamide
40	6-phenyl-3-(propylamino)pyrazine-2-carboxamide
41	3-amino-6-biphenyl-3-ylpyrazine-2-carboxamide
42	3-[5-amino-6-(aminocarbonyl)pyrazin-2-yl]benzoic acid
43	3-amino-6-{4-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide
44	3-amino-N-methyl-6-{3-[(phenylmethyl)oxy]phenyl}pyrazine-2-carboxamide
45	3-amino-6-(3-hydroxyphenyl)-N-methylpyrazine-2-carboxamide
46	3-amino-N-[2-(methyloxy)ethyl]-6-phenylpyrazine-2-carboxamide
47	N-[2-(acetylamino)ethyl]-3-amino-6-phenylpyrazine-2-carboxamide
48	3-amino-6-phenylpyrazine-2-carbohydrazide
49	3-amino-N-hydroxy-6-phenylpyrazine-2-carboxamide
50	3-amino-6-[3-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide
51	3-amino-6-(3-aminophenyl)-N-methylpyrazine-2-carboxamide
52	3-amino-6-[4-(ethyloxy)phenyl]-N-methylpyrazine-2-carboxamide
53	3-amino-N-methyl-6-(3-methylphenyl)pyrazine-2-carboxamide
54	3-amino-N-methyl-6-[4-(methyloxy)phenyl]pyrazine-2-carboxamide
55	3-amino-6-biphenyl-3-yl-N-methylpyrazine-2-carboxamide
56	6-[3-(acetylamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide
57	3-amino-6-[3-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide
58	3-amino-6-[4-(hydroxymethyl)phenyl]-N-methylpyrazine-2-carboxamide
59	3-amino-N-methyl-6-{3-[(1E)-N-methylethanimidoyl]phenyl}pyrazine-2-carboxamide

#	Name
60	3-amino-N-methyl-6-[3-(methyloxy)phenyl]pyrazine-2-carboxamide
61	3-amino-6-(4-fluoro-3-methylphenyl)-N-methylpyrazine-2-carboxamide
62	3-amino-N-methyl-6-pyridin-3-ylpyrazine-2-carboxamide
63	6-[4-(acetylamino)phenyl]-3-amino-N-methylpyrazine-2-carboxamide
64	3-amino-N-methyl-6-[3-[(methylamino)carbonyl]phenyl]pyrazine-2-carboxamide
65	6-(3-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide
66	3-amino-N-methyl-6-(5-pyridin-3-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
67	3-amino-6-(5-cyclopentyl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide
68	3-amino-6-(5-cyclohexyl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide
69	3-amino-N-methyl-6-(5-phenyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
70	3-amino-N-methyl-6-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide
71	3-amino-N-methyl-6-(5-methyl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
72	3-amino-N-methyl-6-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]pyrazine-2-carboxamide
73	3-amino-N-methyl-6-(5-pyridin-4-yl-1,2,4-oxadiazol-3-yl)pyrazine-2-carboxamide
74	3-amino-6-[5-(1,1-dimethylethyl)-1,2,4-oxadiazol-3-yl]-N-methylpyrazine-2-carboxamide
75	3-amino-6-(5-furan-2-yl-1,2,4-oxadiazol-3-yl)-N-methylpyrazine-2-carboxamide
76	3-amino-6-[4-(dimethylamino)phenyl]-N-methylpyrazine-2-carboxamide
77	3-amino-N-methyl-6-[4-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide
78	3-amino-6-(3,5-dimethylphenyl)-N-methylpyrazine-2-carboxamide
79	6-(4-acetylphenyl)-3-amino-N-methylpyrazine-2-carboxamide
80	3-amino-6-[3,4-bis(methyloxy)phenyl]-N-methylpyrazine-2-carboxamide
81	3-amino-N-methyl-6-[3-[(phenylcarbonyl)amino]phenyl]pyrazine-2-carboxamide
82	3-amino-N-methyl-6-[3-[(methylsulfonyl)amino]phenyl]pyrazine-2-carboxamide

#	Name
83	3-amino-N-methyl-6-[3-(1H-tetrazol-5-yl)phenyl]pyrazine-2-carboxamide
84	3-amino-N-methyl-6-[3-(((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
85	3-amino-6-[3-(aminocarbonyl)phenyl]-N-methylpyrazine-2-carboxamide
86	3-amino-6-[3-((dimethylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
87	3-amino-N-methyl-6-[4-(methylsulfonyl)phenyl]pyrazine-2-carboxamide
88	3-amino-6-(3-aminophenyl)-N-ethylpyrazine-2-carboxamide
89	3-amino-6-[3-(((4-fluorophenyl)amino)carbonyl)amino]phenyl]-N-methylpyrazine-2-carboxamide
90	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)amino)carbonyl)amino]phenyl]pyrazine-2-carboxamide
91	6,6'-[(oxomethanediy)bis(iminobenzene-3,1-diyl)]bis(3-amino-N-methylpyrazine-2-carboxamide)
92	3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzoic acid
93	3-amino-N-methyl-6-[3-[(4-methylpiperazin-1-yl)carbonyl]phenyl]pyrazine-2-carboxamide
94	3-amino-N-methyl-6-[3-[(2-phenylethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
95	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
96	3-amino-6-[3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
97	3-amino-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
98	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
99	3-amino-N-methyl-6-[3-([methyl(phenylmethyl)amino]carbonyl)phenyl]pyrazine-2-carboxamide
100	3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
101	3-amino-N-methyl-6-[3-(((4-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
102	3-amino-N-methyl-6-[3-(((4-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
103	3-amino-6-[3-(((4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
104	3-amino-6-[3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
105	3-amino-N-methyl-6-[3-(((3-(methyloxy)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide

#	Name
106	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide
107	3-amino-N-methyl-6-[3-(((3-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
108	3-amino-N-methyl-6-[3-(((1R)-1-[4-(methyloxy)phenyl]ethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
109	3-amino-N-methyl-6-[3-(((2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
110	3-amino-6-[3-(((3-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
111	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
112	3-amino-6-[3-(((1-(4-fluorophenyl)ethyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
113	3-amino-6-[3-(((2,3-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
114	3-amino-6-[3-(((bis(phenylmethyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
115	3-amino-6-[3-(((2,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
116	3-amino-6-[3-(((ethyl(pyridin-4-ylmethyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
117	3-amino-N-methyl-6-[3-(((4-(phenylmethyl)piperazin-1-yl)carbonyl]phenyl)pyrazine-2-carboxamide
118	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)piperidin-4-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
119	3-amino-N-methyl-6-[3-(((4-methylphenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide
120	3-amino-N-methyl-6-[3-(((5-methylpyrazin-2-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
121	3-amino-N-methyl-6-[3-(((pyridin-2-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
122	3-amino-N-methyl-6-[3-(((pyridin-3-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
123	3-amino-N-methyl-6-[3-(((pyridin-4-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
124	3-amino-6-[3-(((furan-2-ylcarbonyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
125	3-amino-6-[3-(((3-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
126	3-amino-6-[3-(((cyclohexylmethyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
127	3-amino-N-methyl-6-[3-(((2-phenylpropyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
128	3-amino-6-[3-(((2-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
129	3-amino-6-[3-(((2-(2-chlorophenyl)ethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
130	3-amino-N-methyl-6-[3-(((2-(2-(methyloxy)phenyl)ethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
131	3-amino-6-[3-(((2-(4-fluorophenyl)ethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
132	3-amino-N-methyl-6-[3-(((2-(2-thienyl)ethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
133	3-amino-6-[3-((cyclohexylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
134	1,1-dimethylethyl 4-(((3-(5-amino-6-((methylamino)carbonyl)pyrazin-2-yl)phenyl)carbonyl)amino)methyl)piperidine-1-carboxylate
135	3-amino-N-methyl-6-[3-((piperidin-4-ylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
136	3-amino-N-methyl-6-[3-(5-phenyl-1,2,4-oxadiazol-3-yl)phenyl]pyrazine-2-carboxamide
137	3-amino-N-methyl-6-[3-[5-(phenylmethyl)-1,2,4-oxadiazol-3-yl]phenyl]pyrazine-2-carboxamide
138	3-amino-N-methyl-6-[3-[5-(2-phenylethyl)-1,2,4-oxadiazol-3-yl]phenyl]pyrazine-2-carboxamide
139	3-amino-6-[3-(((1,1-dimethylethyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
140	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
141	3-amino-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
142	3-amino-N-methyl-6-[3-((phenylamino)carbonyl)phenyl]pyrazine-2-carboxamide
143	3-amino-6-[3-(1,3-dihydro-2H-isoindol-2-ylcarbonyl)phenyl]-N-methylpyrazine-2-carboxamide
144	3-amino-N-methyl-6-[3-(((1R)-1-phenylethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
145	3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
146	3-amino-N-methyl-6-[3-(((1R)-1-phenylpropyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
147	3-amino-N-methyl-6-[3-(((1S)-1-phenylpropyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
148	3-amino-N-methyl-6-[3-(((1R,2S)-2-phenylcyclopropyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
149	phenylmethyl 4-[5-amino-6-((methylamino)carbonyl)pyrazin-2-yl]benzoate
150	3-amino-N-methyl-6-[4-(((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
151	3-amino-N-methyl-6-[4-(((2-phenylethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide

#	Name
152	3-amino-6-[4-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
153	3-amino-N-methyl-6-[4-(((4-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
154	3-amino-6-(4-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
155	3-amino-6-(3-(((4-chlorophenyl)acetyl)amino)phenyl)-N-methylpyrazine-2-carboxamide
156	3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide
157	3-amino-6-[3-((diphenylacetyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
158	3-amino-N-methyl-6-[3-((phenylacetyl)amino)phenyl]pyrazine-2-carboxamide
159	3-amino-N-methyl-6-[3-((3-phenylpropanoyl)amino)phenyl]pyrazine-2-carboxamide
160	3-amino-N-methyl-6-[3-(((1R)-1,2,3,4-tetrahydronaphthalen-1-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide
161	3-amino-N-methyl-6-[3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide
162	3-amino-N-methyl-6-[3-(((3,4,5-tris(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
163	3-amino-N-methyl-6-[3-(((1R,2R)-2-((phenylmethyl)oxy)cyclopentyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
164	3-amino-N-methyl-6-[3-(((1S,2S)-2-((phenylmethyl)oxy)cyclopentyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
165	3-amino-6-(3-cyanophenyl)-N-methylpyrazine-2-carboxamide
166	3-amino-6-[3-((cyclopentylcarbonyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
167	3-amino-6-[3-(((4-cyanophenyl)carbonyl)amino)phenyl]-N-methylpyrazine-2-carboxamide
168	3-amino-N-methyl-6-[3-(((phenylmethyl)amino)carbonyl)amino]phenyl]pyrazine-2-carboxamide
169	3-amino-N-methyl-6-[3-(((naphthalen-2-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
170	3-amino-N-methyl-6-[3-(((tetrahydrofuran-2-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
171	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
172	3-amino-6-[3-(((3-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
173	3-amino-6-[3-(((2-[3,5-bis(methyloxy)phenyl]ethyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
174	3-amino-N-methyl-6-[3-((2-phenylethyl)oxy)phenyl]pyrazine-2-carboxamide

#	Name
175	3-amino-N-methyl-6-(3-([2-(4-methylpiperidin-1-yl)ethyl]oxy)phenyl)pyrazine-2-carboxamide
176	3-amino-6-[3-((2-[4-(2-hydroxyethyl)piperidin-1-yl]ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
177	3-amino-6-(3-([2-(3-hydroxypyrrolidin-1-yl)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
178	3-amino-6-[3-((2-[ethyl(phenylmethyl)amino]ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
179	3-amino-6-(3-([2-(diethylamino)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
180	3-amino-6-(3-(((biphenyl-2-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
181	3-amino-N-methyl-6-(3-([2-(4-phenylpiperazin-1-yl)ethyl]oxy)phenyl)pyrazine-2-carboxamide
182	3-amino-6-(3-(((biphenyl-4-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
183	3-amino-N-methyl-6-[3-((2-[methyl(phenylmethyl)amino]ethyl]oxy)phenyl)pyrazine-2-carboxamide
184	3-amino-N-methyl-6-[3-(((4-(phenyloxy)phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
185	3-amino-N-methyl-6-(3-([2-(morpholin-4-ylethyl)oxy]phenyl)pyrazine-2-carboxamide
186	3-amino-6-[3-((2-[(cyclopropylmethyl)(propyl)amino]ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
187	3-amino-N-methyl-6-[3-((2-[4-(phenylmethyl)piperidin-1-yl]ethyl]oxy)phenyl)pyrazine-2-carboxamide
188	3-amino-N-methyl-6-[3-([2-(piperidin-1-ylethyl)oxy]phenyl)pyrazine-2-carboxamide
189	3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
190	3-amino-6-(3-([2-(4-ethylpiperazin-1-yl)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
191	3-amino-N-methyl-6-[3-([2-(pyrrolidin-1-ylethyl)oxy]phenyl)pyrazine-2-carboxamide
192	3-amino-N-methyl-6-(3-([2-(4-methylpiperazin-1-yl)ethyl]oxy)phenyl)pyrazine-2-carboxamide
193	3-amino-6-[3-((2-[4-(2-hydroxyethyl)piperazin-1-yl]ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
194	3-amino-6-(3-([2-(3,4-dihydroisoquinolin-2(1H)-yl)ethyl]oxy)phenyl)-N-methylpyrazine-2-carboxamide
195	ethyl 4-[2-((3-[5-amino-6-((methylamino)carbonyl)pyrazin-2-yl]phenyl)oxy)ethyl]piperazine-1-carboxylate
196	3-amino-N-methyl-6-[3-(((3S)-1-(phenylmethyl)pyrrolidin-3-yl]amino)carbonyl)phenyl]pyrazine-2-carboxamide
197	3-amino-N-methyl-6-[3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl]amino)carbonyl)phenyl]pyrazine-2-carboxamide

#	Name
198	3-amino-N-methyl-6-[3-([2-(phenyloxy)ethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
199	3-amino-N-methyl-6-[3-([2-[4-(phenylmethyl)piperazin-1-yl]ethyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
200	3-amino-N-methyl-6-[3-([1-(phenylmethyl)piperidin-4-yl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
201	3-amino-6-[3-([furan-3-ylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
202	3-amino-6-[3-([2,3-dihydro-1,4-benzodioxin-2-ylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
203	3-amino-6-[3-([furan-2-ylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
204	3-amino-N-methyl-6-[3-([4-(phenylmethyl)morpholin-2-yl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
205	3-amino-6-[3-([2,3-dihydro-1,4-benzodioxin-6-ylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
206	3-amino-6-[3-([2,3-dihydro-1,4-benzodioxin-6-ylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
207	3-amino-6-[3-([2,3-dihydro-1-benzofuran-5-ylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
208	3-amino-6-[3-([6-fluoro-4H-1,3-benzodioxin-8-yl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
209	3-amino-6-[3-([2,4-dichlorophenyl]acetyl]amino)phenyl]-N-methylpyrazine-2-carboxamide
210	3-amino-6-[3-([3,4-dichlorophenyl]acetyl]amino)phenyl]-N-methylpyrazine-2-carboxamide
211	3-amino-N-methyl-6-[3-([4-(trifluoromethyl)phenyl]acetyl]amino)phenyl]pyrazine-2-carboxamide
212	3-amino-6-[3-([3,5-dimethylphenyl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
213	3-amino-6-[3-([5-chloro-2-methylphenyl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
214	3-amino-6-[3-([2,4-dimethylphenyl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
215	3-amino-6-[3-([2,3-difluorophenyl]methyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
216	3-amino-N-methyl-6-[3-([4-(methylthio)phenyl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
217	3-amino-N-methyl-6-[3-([2-methylpropyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
218	3-amino-N-methyl-6-[3-([3-[(trifluoromethyl)oxy]phenyl]methyl]amino)carbonyl]phenyl]pyrazine-2-carboxamide
219	3-amino-6-[3-([cyclopropylmethyl]amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
220	3-amino-6-[3-([cyclopentylamino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
221	3-amino-N-methyl-6-(3-(((4-methylphenyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
222	3-amino-N-methyl-6-[3-(((4-(methyloxy)phenyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
223	3-amino-6-(3-(((1,3-benzodioxol-5-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
224	3-amino-6-(3-(((biphenyl-3-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
225	3-amino-N-methyl-6-(3-(((1-methyl-1-phenylethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
226	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
227	3-amino-N-methyl-6-(3-(((4-((trifluoromethyl)oxy)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
228	3-amino-6-[3-(((4-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
229	3-amino-6-[3-(((4-chloro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
230	3-amino-6-[3-(((4-aminophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
231	3-amino-6-[3-(((2-aminophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
232	3-amino-6-[3-(((3-aminophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
233	3-amino-N-methyl-6-pyrimidin-5-ylpyrazine-2-carboxamide
234	3-amino-N-methyl-6-[3-(methylsulfonyl)phenyl]pyrazine-2-carboxamide
235	3-amino-N-methyl-6-[4-(tetrahydro-2H-pyran-2-yloxy)phenyl]pyrazine-2-carboxamide
236	3-amino-6-(4-hydroxyphenyl)-N-methylpyrazine-2-carboxamide
237	3-amino-6-[3-(((4-(aminosulfonyl)phenyl)methyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
238	3-amino-6-[3-(((4-(dimethylamino)phenyl)methyl)amino)carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
239	3-amino-6-[3-(((3-bromophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
240	3-amino-6-(4-chloro-3-((phenylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
241	methyl 3-{5-amino-6-[(cyclopropylamino)carbonyl]pyrazin-2-yl}benzoate
242	methyl 3-{5-amino-6-[(ethylamino)carbonyl]pyrazin-2-yl}benzoate

#	Name
243	3-amino-6-[3-(((2,3-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
244	3-amino-6-[3-(((4-hydroxy-3-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
245	3-amino-6-[3-(((4-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
246	3-amino-6-[3-(((2-(ethyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
247	3-amino-6-[3-(((2-chloro-6-(phenyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
248	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
249	3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
250	3-amino-6-[3-(((2-chloro-6-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
251	3-amino-N-methyl-6-[3-(((2,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
252	3-amino-6-[3-(((2,2-dimethylpropyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
253	3-amino-6-[3-(((cyclopentyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
254	3-amino-6-[3-(((3,5-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
255	3-amino-N-methyl-6-[3-(((4-(methylsulfonyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
256	3-amino-6-[3-(((5-bromo-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
257	3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
258	3-amino-N-methyl-6-[3-(((2,4,6-tris(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
259	3-amino-N-methyl-6-[3-(((3-methylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
260	3-amino-6-[3-(((4-bromo-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
261	3-amino-6-[3-(((2,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
262	3-amino-6-[3-(((3,5-bis(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
263	3-amino-6-[3-(((2,2-diphenylethyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
264	3-amino-N-methyl-6-(3-thienyl)pyrazine-2-carboxamide
265	3-amino-N-methyl-6-[3-(((phenylmethyl)amino)sulfonyl]phenyl]pyrazine-2-carboxamide

#	Name
266	5,5'-diamino-N,N'-dimethyl-2,2'-bipyrazine-6,6'-dicarboxamide
267	3-amino-6-(1H-indol-5-yl)-N-methylpyrazine-2-carboxamide
268	3-amino-N-methyl-6-(2-thienyl)pyrazine-2-carboxamide
269	3-amino-N-methyl-6-pyridin-4-ylpyrazine-2-carboxamide
270	3-amino-N-cyclopropyl-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
271	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-cyclopropylpyrazine-2-carboxamide
272	3-amino-N-ethyl-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
273	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-ethylpyrazine-2-carboxamide
274	3-amino-6-[3-(((2,3-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
275	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
276	3-amino-6-[3-(((3,5-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
277	3-amino-6-[3-(((5-chloro-2-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
278	3-amino-6-[3-((cyclopropylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
279	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
280	3-amino-6-[3-(((4'-fluorobiphenyl-2-yl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
281	3-amino-N-methyl-6-[3-(((2,3,4-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
282	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
283	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
284	3-amino-6-[3-((9H-fluoren-9-ylamino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
285	3-amino-N-methyl-6-[3-(((2-methylphenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
286	3-amino-6-[3-(((3,4-dimethylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
287	3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide
288	3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl)phenyl]-N-methylpyrazine-2-carboxamide

#	Name
289	3-amino-6-[3-(((2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
290	3-amino-N-methyl-6-[3-(((4-(4-methylpiperazin-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
291	3-amino-6-(4-fluoro-3-(((phenylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
292	3-amino-N-methyl-6-[3-(((2,4,6-trichlorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
293	3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
294	3-amino-6-[3-(((2,5-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
295	3-amino-6-[3-(((3-fluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
296	3-amino-6-[3-(((2-chloro-6-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
297	3-amino-6-[3-(((2-chloro-3,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
298	3-amino-6-[3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
299	3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
300	3-amino-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
301	3-amino-6-[3-(((6-chloro-2-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
302	3-amino-6-[3-(((4-iodophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
303	3-amino-6-[3-(((3-chloro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
304	3-amino-6-[3-(((4-bromophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
305	3-amino-6-[3-(((2-bromophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
306	3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
307	3-amino-N-methyl-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
308	3-amino-N-methyl-6-(3-(((phenylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
309	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
310	3-amino-6-[3-(((2,5-difluorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
311	3-amino-N-methyl-6-(3-(((naphthalen-2-ylcarbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide

#	Name
312	3-amino-N-methyl-6-(3-[(phenylmethyl)thio]methyl)phenyl)pyrazine-2-carboxamide
313	3-amino-N-methyl-6-(3-[(phenylmethyl)sulfonyl]methyl)phenyl)pyrazine-2-carboxamide
314	3-amino-N-methyl-6-(3-[(phenylmethyl)oxy]methyl)phenyl)pyrazine-2-carboxamide
315	3-amino-N-methyl-6-(3-[(phenylmethyl)amino]methyl)phenyl)pyrazine-2-carboxamide
316	3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
317	9H-fluoren-9-ylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
318	phenylmethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
319	ethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
320	3-amino-6-[3-(((3,4-dichlorophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
321	1,1-dimethylethyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
322	(1R,2S,5R)-5-methyl-2-(1-methylethyl)cyclohexyl (3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbamate
323	3-amino-6-[3-(((4-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
324	3-amino-N-methyl-6-(3-(((4-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
325	3-amino-N-methyl-6-[3-(((2-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
326	3-amino-6-[3-(((3-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
327	3-amino-6-[3-(((2-hydroxyphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
328	3-amino-N-methyl-6-[3-((2-phenylhydrazino)carbonyl)phenyl]pyrazine-2-carboxamide
329	3-amino-N-(2-hydroxyethyl)-6-(3-methylphenyl)pyrazine-2-carboxamide
330	3-amino-N-methyl-6-(3-[(phenylacetyl)amino]methyl)phenyl)pyrazine-2-carboxamide
331	3-amino-6-[3-(((4-chlorophenyl)acetyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
332	3-amino-6-(3-(((biphenyl-4-ylcarbonyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
333	3-amino-6-[3-(((2-chloro-5-(trifluoromethyl)phenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
334	N-[(3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)methyl]-1H-indole-2-carboxamide

#	Name
335	3-amino-6-[3-(((3-hydroxypyridin-2-yl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
336	3-amino-6-(3-(((1H-indol-3-ylacetyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
337	3-amino-6-[3-(((2,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
338	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
339	3-amino-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
340	3-amino-6-[3-(((3-iodophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
341	3-amino-N-methyl-6-[3-(((2-morpholin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
342	3-amino-N-methyl-6-(3-(((3-[(2-morpholin-4-ylethyl)oxy]phenyl)methyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
343	3-amino-N-methyl-6-[3-(((4-morpholin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
344	3-amino-N-methyl-6-[3-(((3-morpholin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
345	3-amino-N-methyl-6-[3-(((4-(1-methylethyl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
346	3-amino-N-methyl-6-(3-(((4-(1-methylethyl)phenyl)oxy)acetyl)amino)methyl]phenyl]pyrazine-2-carboxamide
347	3-amino-6-[3-(((4-aminophenyl)thio)acetyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
348	3-amino-N-methyl-6-(4-(methyloxy)-3-(((phenylmethyl)amino)carbonyl]phenyl)pyrazine-2-carboxamide
349	3-amino-N-methyl-6-(5-(((phenylmethyl)amino)carbonyl)pyridin-3-yl)pyrazine-2-carboxamide
350	3-amino-6-[3-(((4-iodophenyl)carbonyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
351	3-amino-N-methyl-6-[3-(((4-pentylphenyl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide
352	3-amino-6-(3-(((1,3-benzodioxol-5-ylcarbonyl)amino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
353	3-amino-6-[3-(((4-(1,1-dimethylethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
354	3-amino-N-methyl-6-(3-(((1-[5-(trifluoromethyl)pyrimidin-2-yl]piperidin-4-yl)carbonyl)amino)methyl]phenyl)pyrazine-2-carboxamide
355	3-amino-N-methyl-6-[3-(((2-piperidin-1-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
356	3-amino-N-methyl-6-[3-(((2-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
357	1,1-dimethylethyl (3R)-3-(((3-{5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl}phenyl)carbonyl)amino)pyrrolidine-1-carboxylate

#	Name
358	3-amino-N-methyl-6-(3-(((3R)-pyrrolidin-3-ylamino)carbonyl)phenyl)pyrazine-2-carboxamide
359	3-(((4-chlorophenyl)methyl)amino)-6-(3-(((3R)-1-((4-chlorophenyl)methyl)pyrrolidin-3-yl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
360	3-amino-6-(3-(((3R)-1-((2,4-dichlorophenyl)methyl)pyrrolidin-3-yl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
361	N-methyl-3-(((3-(methyloxy)phenyl)methyl)amino)-6-(3-(((3R)-1-((3-(methyloxy)phenyl)methyl)pyrrolidin-3-yl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
362	3-amino-6-(3-(((3R)-1-((2,6-dichlorophenyl)methyl)pyrrolidin-3-yl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
363	N-methyl-3-((phenylmethyl)amino)-6-(3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
364	1,1-dimethylethyl 3-(((3-(5-amino-6-((methylamino)carbonyl)pyrazin-2-yl)phenyl)carbonyl)amino)methyl)pyrrolidine-1-carboxylate
365	1,1-dimethylethyl 2-(((3-(5-amino-6-((methylamino)carbonyl)pyrazin-2-yl)phenyl)carbonyl)amino)methyl)pyrrolidine-1-carboxylate
366	3-amino-N-methyl-6-(3-(((pyrrolidin-3-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
367	3-amino-N-methyl-6-(3-(((pyrrolidin-2-ylmethyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
368	3-amino-6-(2-fluoro-5-(((phenylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
369	3-amino-6-(3-((1,3-benzodioxol-5-ylacetyl)amino)phenyl)-N-methylpyrazine-2-carboxamide
370	3-amino-6-(3-(((2-chloro-4-fluorophenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
371	3-amino-6-(3-(((3,4-bis(methyloxy)phenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
372	3-amino-N-methyl-6-(3-((phenylmethyl)amino)phenyl)pyrazine-2-carboxamide
373	3-amino-N-methyl-6-(3-(((2-((2-morpholin-4-ylethyl)oxy)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
374	3-amino-N-methyl-6-(3-(((4-(1-methylethyl)phenyl)carbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
375	3-amino-6-(3-(((4-butylphenyl)methyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide
376	3-amino-N-methyl-6-(3-(((4-(1,2,3-thiadiazol-4-yl)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
377	3-amino-N-methyl-6-(3-(((4-(2-thienyl)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
378	3-amino-6-(3-(((4-(1,1-dimethylethyl)phenyl)carbonyl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
379	3-amino-N-methyl-6-(3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl)phenyl)pyrazine-2-carboxamide
380	3-amino-6-(3-(((3,4-dihydro-2H-1,4-benzoxazin-6-ylmethyl)amino)carbonyl)phenyl)-N-methylpyrazine-2-carboxamide

#	Name
381	3-amino-6-[3-(((2-amino-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
382	3-amino-6-[3-(((biphenyl-4-ylmethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
383	3-amino-6-[3-(((biphenyl-2-ylmethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
384	3-amino-6-[3-(((biphenyl-3-ylmethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
385	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)pyrrolidin-2-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
386	3-amino-6-[3-(((3,4-dihydro-2H-1,4-benzoxazin-6-ylmethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
387	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
388	3-amino-6-[3-(((2,5-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
389	3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
390	3-amino-N-methyl-6-[3-(((2-pyridin-4-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
391	3-amino-N-methyl-6-[3-(((2-pyridin-3-ylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
392	3-amino-N-methyl-6-[3-(((4-(methylsulfonyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
393	3-amino-6-[3-(((2,2-diphenylethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
394	3-amino-N-methyl-6-[3-(((1-(phenylmethyl)pyrrolidin-3-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
395	3-amino-N-methyl-6-[3-(((3-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
396	3-amino-6-[3-(((4-aminophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
397	3-amino-N-methyl-6-[3-(((4-(phenyloxy)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
398	3-amino-6-[3-(((2,6-bis(methyloxy)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
399	3-amino-6-[3-(((2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
400	3-amino-6-[5-(((2,6-difluorophenyl)methyl)amino)carbonyl]pyridin-3-yl]-N-methylpyrazine-2-carboxamide
401	3-amino-6-[5-(((biphenyl-2-ylmethyl)amino)carbonyl]pyridin-3-yl]-N-methylpyrazine-2-carboxamide
402	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
403	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide

#	Name
404	3-amino-N-methyl-6-[3-(((2-piperidin-1-yl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
405	3-amino-6-[3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
406	3-amino-N-methyl-6-[3-(((6-(trifluoromethyl)pyridin-3-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
407	3-amino-6-[3-(1H-benzimidazol-2-yl)phenyl]-N-methylpyrazine-2-carboxamide
408	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
409	3-amino-6-[3-(((3,5-dichlorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
410	3-amino-N-methyl-6-[3-((2-phenylethyl)amino)phenyl]pyrazine-2-carboxamide
411	3-amino-6-[5-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]pyridin-3-yl]-N-methylpyrazine-2-carboxamide
412	3-amino-N-methyl-6-[3-(((2-(phenylamino)ethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
413	3-amino-6-[3-(((4-ethylphenyl)methyl)amino)carbonyl]phenyl]-N-methylpyrazine-2-carboxamide
414	3-amino-N-methyl-6-[3-(((4-pyridin-3-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
415	3-amino-6-[3-((2,3-dihydro-1H-inden-2-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
416	3-amino-6-[3-(1,3-dihydro-2H-isoindol-2-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide
417	3-amino-6-[3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
418	3-amino-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
419	3-amino-N-methyl-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
420	3-amino-N-methyl-6-[3-(((1R)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide
421	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
422	3-amino-N-methyl-6-[3-(((4-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
423	3-amino-N-methyl-6-[3-(((3-(1,2,3-thiadiazol-4-yl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
424	3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
425	3-amino-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
426	3-amino-N-methyl-6-[3-(((3-(trifluoromethyl)phenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide

#	Name
427	3-amino-N-methyl-6-[3-(((1S)-1-phenylethyl)amino)methyl]phenyl]pyrazine-2-carboxamide
428	3-amino-6-[3-(((1R)-2-hydroxy-1-phenylethyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
429	methyl (2S)-(((3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)phenyl)methyl]amino)(phenyl)ethanoate
430	3-amino-6-(3-[[2-(2,5-difluorophenyl)hydrazino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
431	3-amino-N-methyl-6-(3-[[2-(2,3,5,6-tetrafluoro-4-methylphenyl)hydrazino]carbonyl]phenyl)pyrazine-2-carboxamide
432	3-amino-6-(3-[[2-(2-fluorophenyl)hydrazino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
433	3-amino-6-(3-[[2-(2,4-difluorophenyl)hydrazino]carbonyl]phenyl)-N-methylpyrazine-2-carboxamide
434	3-amino-N-methyl-6-(3-[[2-(2,3,5,6-tetrafluorophenyl)hydrazino]carbonyl]phenyl)pyrazine-2-carboxamide
435	3-amino-N-methyl-6-[3-(((5-methylpyrazin-2-yl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
436	3-amino-N-methyl-6-(3-(((1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
437	3-amino-N-methyl-6-[3-(((3S)-1-(phenylmethyl)pyrrolidin-3-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
438	3-amino-N-methyl-6-[3-(((3R)-1-(phenylmethyl)pyrrolidin-3-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
439	3-amino-6-(3-[[ethyl(phenylmethyl)amino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
440	3-amino-N-methyl-6-[3-(((1S,2S)-2-phenylcyclopropyl)amino)methyl]phenyl]pyrazine-2-carboxamide
441	3-Amino-6-(3-benzylcarbamoyl-phenyl)-pyrazine-2-carboxylic acid methylamide
442	3-amino-N-methyl-6-[3-(4-phenyl-1H-imidazol-2-yl)phenyl]pyrazine-2-carboxamide
443	3-amino-N-methyl-6-[3-(((4-propylphenyl)methyl)amino)methyl]phenyl]pyrazine-2-carboxamide
444	3-amino-6-[3-(((5-bromo-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
445	3-amino-N-methyl-6-(3-(((2R)-1,2,3,4-tetrahydronaphthalen-2-ylamino)methyl)phenyl)pyrazine-2-carboxamide
446	3-amino-N-methyl-6-(3-(((1-methyl-1-phenylethyl)amino)methyl)phenyl)pyrazine-2-carboxamide
447	3-amino-6-(3-[(9H-fluoren-9-ylamino)methyl]phenyl)-N-methylpyrazine-2-carboxamide
448	3-amino-N-methyl-6-[3-[(naphthalen-1-ylamino)methyl]phenyl]pyrazine-2-carboxamide
449	3-amino-6-[3-(((3-fluorobiphenyl-4-yl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide

#	Name
450	3-amino-6-[3-(((2-fluoro-4-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
451	3-amino-6-[3-(((2-fluoro-4-furan-2-ylphenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
452	3-amino-N-methyl-6-[3-((naphthalen-2-ylamino)methyl)phenyl]pyrazine-2-carboxamide
453	3-amino-6-[3-(((1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
454	3-amino-N-methyl-6-[3-(((2S)-1,2,3,4-tetrahydronaphthalen-2-ylamino)methyl)phenyl]pyrazine-2-carboxamide
455	methyl 3-amino-6-[3-(((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxylate
456	3-amino-6-[3-(((5-bromo-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
457	3-amino-6-[3-(((2-fluoro-5-(2-thienyl)phenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
458	3-amino-6-[3-(((4-ethenyl-2-fluorophenyl)methyl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
459	3-amino-6-[3-(((4,7-difluoro-3-methyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
460	3-amino-N-methyl-6-[3-(((1S,2R)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
461	3-amino-6-[3-(((5-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
462	3-amino-6-[3-(((2S)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
463	3-amino-6-[3-(((2R)-6-bromo-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
464	3-amino-6-[3-(((5-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
465	3-amino-6-[3-(((2S)-5-bromo-2,3-dihydro-1H-inden-2-yl)amino)methyl]phenyl]-N-methylpyrazine-2-carboxamide
466	3-amino-6-[3-(((4-fluoro-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
467	3-amino-6-[3-(((4-fluoro-1-hydroxy-2,3-dihydro-1H-inden-2-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
468	3-amino-N-methyl-6-[3-(((1R,2S)-2-(methyloxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
469	3-amino-6-[3-(((4,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
470	3-amino-6-[3-(((6-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
471	3-amino-N-methyl-6-[3-(((3-oxo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
472	3-amino-6-[3-(((5-furan-2-yl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide

#	Name
473	3-amino-6-(3-(((4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
474	3-amino-6-(3-(((4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
475	3-amino-6-(3-(((6-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
476	3-amino-6-(3-(((5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
477	3-amino-6-(3-(((5-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
478	3-amino-N-methyl-6-[3-(((5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
479	3-amino-6-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
480	3-amino-6-(3-(((5-furan-3-yl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
481	3-amino-N-methyl-6-[3-(((5-(3-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
482	3-amino-6-[3-(((1S)-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
483	3-amino-6-[3-(((1S)-5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
484	3-amino-6-(3-(((7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
485	3-amino-6-(3-(((6-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
486	3-amino-6-(3-(((4,6-dichloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
487	3-amino-6-(3-(((4-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
488	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-ethylpyrazine-2-carboxamide
489	3-amino-6-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-ethylpyrazine-2-carboxamide
490	3-amino-N-methyl-6-[3-(((1S)-5-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
491	3-amino-N-methyl-6-(3-(((5-phenyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)pyrazine-2-carboxamide
492	3-amino-6-(3-(((4-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
493	3-amino-N-methyl-6-(3-(((4-phenyl-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)pyrazine-2-carboxamide
494	3-amino-N-methyl-6-[3-(((4-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
495	3-amino-N-methyl-6-[3-(((6-(2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide

#	Name
496	3-amino-N-methyl-6-[3-(((5-(4-methyl-2-thienyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
497	3-amino-N-methyl-6-[3-(((2R)-6-phenyl-1,2,3,4-tetrahydronaphthalen-2-yl)amino)methyl]phenyl]pyrazine-2-carboxamide
498	3-amino-N-methyl-6-naphthalen-2-ylpyrazine-2-carboxamide
499	3-amino-N-methyl-6-[3-(((1S,2S)-2-((phenylmethyl)oxy)cyclopentyl)amino)methyl]phenyl]pyrazine-2-carboxamide
500	3-amino-6-[3-((cyclopentylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
501	3-amino-6-[3-(((6-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
502	3-amino-6-[3-(((7-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
503	3-amino-N-methyl-6-quinolin-3-ylpyrazine-2-carboxamide
504	3-amino-6-[3-(1H-imidazol-1-ylmethyl)phenyl]-N-methylpyrazine-2-carboxamide
505	3-amino-N-methyl-6-[3-(((5-((2-morpholin-4-ylethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
506	3-amino-N-methyl-6-[3-(((1R,2R)-2-((phenylmethyl)oxy)cyclopentyl)amino)methyl]phenyl]pyrazine-2-carboxamide
507	3-amino-6-[3-((2,3-dihydro-1-benzofuran-3-ylamino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
508	3-amino-6-[3-(((5-((cyanomethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
509	2-amino-5-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]-N-methylpyridine-3-carboxamide
510	3-amino-N-methyl-6-[3-(((5-(4-methylphenyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
511	3-amino-6-[3-(((2-(dimethylamino)-1-phenylethyl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
512	3-amino-N-methyl-6-[3-(((5-(1H-1,2,4-triazol-1-yl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
513	3-amino-6-[3-(((5-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
514	3-amino-N-methyl-6-[3-(((4-((2-morpholin-4-ylethyl)oxy)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]pyrazine-2-carboxamide
515	3-amino-6-[3-(((4-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl]-N-methylpyrazine-2-carboxamide
516	3-amino-N-methyl-6-[1-((phenylmethyl)amino)isoquinolin-7-yl]pyrazine-2-carboxamide
517	3-amino-N-methyl-6-[3-(((2-morpholin-4-yl-1-phenylethyl)amino)methyl)phenyl]pyrazine-2-carboxamide
518	3-amino-N-methyl-6-[3-(((1-phenyl-2-pyrrolidin-1-ylethyl)amino)methyl)phenyl]pyrazine-2-carboxamide

#	Name
519	2-amino-5-(3-(((6-bromo-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
520	2-amino-5-(3-(((4,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
521	2-amino-5-(3-(((7-bromo-4-fluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
522	2-amino-5-(3-(((4-chloro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
523	2-amino-5-(3-(((5,6-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
524	2-amino-5-(3-(((5,7-difluoro-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyridine-3-carboxamide
525	3-amino-6-(1-((1S)-2,3-dihydro-1H-inden-1-ylamino)isoquinolin-7-yl)-N-methylpyrazine-2-carboxamide
526	3-amino-6-(3-((1S)-2,3-dihydro-1H-inden-1-ylamino)-2,3-dihydro-1H-inden-5-yl)-N-methylpyrazine-2-carboxamide
527	3-amino-6-(3-(((6-(3-hydroxypropyl)-2,3-dihydro-1H-inden-1-yl)amino)methyl)phenyl)-N-methylpyrazine-2-carboxamide
528	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(1,3-oxazol-5-yl)pyrazin-2-amine
529	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carbohydrazide
530	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
531	1-(3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazin-2-yl)ethanone
532	3-amino-6-(3-((2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(naphthalen-2-ylmethyl)pyrazine-2-carboxamide
533	3-amino-N-cyclohexyl-6-(3-((2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
534	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(pyridin-4-ylmethyl)pyrazine-2-carboxamide
535	{3-[5-amino-6-(4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl)methanol
536	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(4H-1,2,4-triazol-3-yl)pyrazin-2-amine
537	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2,2,2-trifluoroethyl)pyrazine-2-carboxamide
538	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(5-hydroxypentyl)pyrazine-2-carboxamide
539	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(4-hydroxybutyl)pyrazine-2-carboxamide
540	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-pyrrolidin-1-ylethyl)pyrazine-2-carboxamide
541	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-piperidin-1-ylethyl)pyrazine-2-carboxamide

#	Name
542	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-morpholin-4-ylethyl)pyrazine-2-carboxamide
543	3-amino-N-(cyclopropylmethyl)-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
544	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(phenylmethyl)pyrazine-2-carboxamide
545	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-phenylethyl)pyrazine-2-carboxamide
546	[3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazin-2-yl](phenyl)methanone
547	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(2-hydroxyethyl)pyrazine-2-carboxamide
548	3-amino-N-cyclopropyl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
549	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(pyrrolidin-1-ylcarbonyl)pyrazin-2-amine
550	{3-[5-amino-6-(5-methyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol
551	{3-[5-amino-6-(5-phenyl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl}methanol
552	(3-{5-amino-6-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl}phenyl)methanol
553	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N'-methylpyrazine-2-carbohydrazide
554	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N,N-dimethylpyrazine-2-carboxamide
555	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(phenylmethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
556	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(dimethylamino)ethyl]pyrazine-2-carboxamide
557	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(3-phenyl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
558	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide
559	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[2-(methylamino)ethyl]pyrazine-2-carboxamide
560	3-{5-amino-6-[(3-fluorophenyl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
561	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-(3-hydroxypropyl)pyrazine-2-carboxamide
562	1,1-dimethylethyl 4-(((3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
563	1,1-dimethylethyl 4-(((3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)methylpiperidine-1-carboxylate
564	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-piperidin-4-ylpyrazine-2-carboxamide

#	Name
565	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-(piperidin-4-ylmethyl)pyrazine-2-carboxamide
566	3-[5-amino-6-(morpholin-4-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
567	1,1-dimethylethyl 3-(((3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
568	3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)-N-pyrrolidin-3-ylpyrazine-2-carboxamide
569	(3-(5-amino-6-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)phenyl)methanol
570	{3-[5-amino-6-(5-furan-2-yl-4H-1,2,4-triazol-3-yl)pyrazin-2-yl]phenyl)methanol
571	[3-(5-amino-6-[5-(3-(methyloxy)phenyl]-4H-1,2,4-triazol-3-yl)pyrazin-2-yl)phenyl)methanol
572	2-((3-amino-6-[3-(hydroxymethyl)phenyl]pyrazin-2-yl)carbonyl)-N-phenylhydrazinecarboxamide
573	3-(5-amino-6-[(4-methylpiperazin-1-yl)carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
574	3-(5-amino-6-[[4-(4-fluorophenyl)piperazin-1-yl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
575	3-(5-amino-6-[[4-(phenylmethyl)piperazin-1-yl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
576	methyl N-[[3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]glycinate
577	N-[[3-amino-6-(3-(((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]glycine
578	3-(5-amino-6-[(3,5-difluorophenyl)carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
579	3-[5-amino-6-(biphenyl-4-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
580	{3-[5-amino-6-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl)methanol
581	(3-[5-amino-6-[3-(4-chlorophenyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl]phenyl)methanol
582	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[5-(1,1-dimethylethyl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
583	(3-[5-amino-6-[3-(2-thienyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl]phenyl)methanol
584	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-[3-(3-(methyloxy)phenyl)-1H-1,2,4-triazol-5-yl]pyrazin-2-amine
585	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[3-(dimethylamino)propyl]pyrazine-2-carboxamide
586	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-N-[3-(1H-imidazol-1-yl)propyl]pyrazine-2-carboxamide
587	3-(5-amino-6-[(4-chloro-3-fluorophenyl)carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide

#	Name
588	3-(5-amino-6-[[2,4-bis(methoxy)phenyl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
589	3-(5-amino-6-[[4-(dimethylamino)phenyl]carbonyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
590	methyl 1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]-L-prolinate
591	methyl 1-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]piperidine-3-carboxylate
592	1,1-dimethylethyl 4-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]piperazine-1-carboxylate
593	3-[5-amino-6-(piperazin-1-ylcarbonyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
594	N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]glutamic acid
595	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-{5-[[methoxy)methyl]-4H-1,2,4-triazol-3-yl}pyrazin-2-amine
596	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-pyridin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
597	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[3-(methylamino)propyl]pyrazine-2-carboxamide
598	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-(3-pyrrolidin-1-ylpropyl)pyrazine-2-carboxamide
599	3-(5-amino-6-[(1E)-N-[4-(methoxy)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
600	3-(5-amino-6-[(1E)-N-[4-(1-methylethyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl)-N-(phenylmethyl)benzamide
601	3-[5-amino-6-[(1E)-N-1,3-benzothiazol-2-ylethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
602	3-[5-amino-6-[(1E)-N-(4-methylphenyl)ethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
603	3-[5-amino-6-[(1E)-N-(4-chlorophenyl)ethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
604	3-[5-amino-6-[(1E)-N-methyl-N-phenylethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
605	3-[5-amino-6-[(1E)-N-(2-hydroxyethyl)ethanehydrazonoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
606	{3-[5-amino-6-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
607	{3-[5-amino-6-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
608	3-[5-amino-6-(2-aminopyrimidin-4-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
609	3-[5-amino-6-[2-(dimethylamino)pyrimidin-4-yl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
610	ethyl N-[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]-L-tyrosinate

#	Name
611	N-[[3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]-D-tyrosine
612	1,1-dimethylethyl [3-((3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino]propyl]carbamate
613	3-amino-N-(3-aminopropyl)-6-(3-((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
614	1-[[3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]piperidine-3-carboxylic
615	1,1-dimethylethyl [2-((3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino]ethyl]carbamate
616	3-amino-N-(2-aminoethyl)-6-(3-((phenylmethyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
617	1,1-dimethylethyl (1-[[3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]carbonyl]piperidin-4-yl)carbamate
618	3-{5-amino-6-[(4-aminopiperidin-1-yl)carbonyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
619	3-{5-amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
620	3-{5-amino-6-[imino(morpholin-4-ylamino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
621	3-{5-amino-6-[imino[(4-methylpiperazin-1-yl)amino]methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
622	3-{5-amino-6-[imino(piperidin-1-ylamino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
623	3-{5-amino-6-[(azepan-1-ylamino)(imino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
624	3-{5-amino-6-[imino(((2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl)amino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
625	{3-[5-amino-6-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
626	3-{5-amino-6-[(1E)-N-morpholin-4-ylethanimidoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
627	4-((2E)-2-{1-[3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]ethylidene}hydrazino)benzoic acid
628	ethyl ((2E)-2-{1-[3-amino-6-(3-((phenylmethyl)amino)carbonyl)phenyl)pyrazin-2-yl]ethylidene}hydrazino)acetate
629	3-{5-amino-6-[(1E)-N,N-dimethylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
630	3-{5-amino-6-[(1E)-N-[4-(methylsulfonyl)phenyl]ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
631	3-{5-amino-6-[(1E)-N-(4-cyanophenyl)ethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
632	3-{5-amino-6-[(1E)-N-pyridin-2-ylethanehydrazonoyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide

#	Name
633	3-(5-amino-6-((1E)-N-[amino(imino)methyl]ethanehydrazonoyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
634	3-[5-amino-6-((1E)-N-[4-((trifluoromethyl)oxy)phenyl]ethanehydrazonoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
635	3-(5-Amino-6-{1-[(4-nitro-phenyl)-hydrazono]-ethyl}-pyrazin-2-yl)-N-benzylbenzamide
636	3-(5-amino-6-((1E)-N-[4-(trifluoromethyl)pyrimidin-2-yl]ethanehydrazonoyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
637	3-[5-amino-6-((1E)-N-1H-1,2,3-benzotriazol-1-ylethanimidoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
638	3-[5-amino-6-((1E)-N-methylethanehydrazonoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
639	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-piperidin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
640	3-(5-amino-6-((1E)-N-[4-(trifluoromethyl)phenyl]ethanehydrazonoyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
641	3-[5-amino-6-((1E)-N-phenylethanehydrazonoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
642	3-[5-amino-6-((1E)-N-(4-methylpiperazin-1-yl)ethanimidoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
643	{3-[5-amino-6-(3-pyrrolidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
644	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
645	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-yl]pyrazine-2-carboxamide
646	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(furan-2-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide
647	N-[3-(acetylamino)propyl]-3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
648	3-amino-N-[3-[(furan-2-ylcarbonyl)amino]propyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
649	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-(pyridin-4-ylcarbonyl)pyrrolidin-3-yl]pyrazine-2-carboxamide
650	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(pyridin-4-ylcarbonyl)amino]propyl]pyrazine-2-carboxamide
651	N-[3-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]propyl]quinoxaline-2-carboxamide
652	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[[[6-(trifluoromethyl)pyridin-3-yl]carbonyl]amino]propyl]pyrazine-2-carboxamide
653	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(quinolin-8-ylsulfonyl)amino]propyl]pyrazine-2-carboxamide
654	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[3-[(pyridin-2-ylsulfonyl)amino]propyl]pyrazine-2-carboxamide
655	3-amino-N-[3-[[[3,5-dimethylisoxazol-4-yl]amino]carbonyl]amino]propyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide

#	Name
656	3-[5-amino-6-(2-methylpyrimidin-4-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
657	3-[5-amino-6-[2-(methyldamino)pyrimidin-4-yl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
658	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[2-[(pyridin-4-ylcarbonyl)amino]ethyl]pyrazine-2-carboxamide
659	3-amino-N-[2-[(furan-2-ylcarbonyl)amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
660	N,N'-cyclohexane-1,2-diylbis[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide]
661	N-[2-(acetylamino)ethyl]-3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
662	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[2-[(quinolin-8-ylsulfonyl)amino]ethyl]pyrazine-2-carboxamide
663	N-[2-[[[3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]ethyl]quinoxaline-2-carboxamide
664	3-amino-N-[2-[[[2-chloropyridin-3-yl]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
665	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]-N-[2-[(pyridin-2-ylsulfonyl)amino]ethyl]pyrazine-2-carboxamide
666	3-amino-N-[2-[[[3,5-dimethylisoxazol-4-yl]amino]carbonyl]amino]ethyl]-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
667	3-amino-6-(3-[[[(phenylmethyl)amino]carbonyl]phenyl]pyrazine-2-carboxamide
668	3-[5-amino-6-(imino{2-[4-(trifluoromethyl)pyrimidin-2-yl]hydrazino}methyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
669	3-[5-amino-6-[2-(1,3-benzothiazol-2-yl)hydrazino](imino)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
670	3-[5-amino-6-(1,5-diphenyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
671	3-[5-amino-6-(1,5-dimethyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
672	3-[5-amino-6-(1-methyl-5-phenyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
673	3-[5-amino-6-((1E)-N-[(2R)-2-[(methyloxy)methyl]pyrrolidin-1-yl]ethanimidoyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
674	3-[5-amino-6-[(1E)-N-azepan-1-ylethanimidoyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
675	3-[5-amino-6-((E)-[(phenylmethyl)hydrazono]methyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
676	3-[5-amino-6-((E)-[amino(imino)methyl]hydrazono)methyl]pyrazin-2-yl]-N-(phenylmethyl)benzamide
677	3-[5-amino-6-((E)-[(2-hydroxyethyl)hydrazono]methyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
678	3-[5-amino-6-((E)-(pyridin-2-ylhydrazono)methyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide

#	Name
679	3-(5-amino-6-((E)-[(4-cyanophenyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
680	3-(5-amino-6-((E)-[(4-methylphenyl)hydrazono]methyl)pyrazin-2-yl)-N-(phenylmethyl)benzamide
681	3-{5-amino-6-[(E)-(hydroxyimino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
682	{3-[5-amino-6-(3-pyridin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
683	5-(3-[[[(1R)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-piperidin-3-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine
684	3-amino-N-[3-[[[4-(dimethylamino)phenyl]carbonyl]amino]propyl]-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
685	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
686	1,1-dimethylethyl 4-[(3S)-3-[[[3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidin-1-yl]piperidine-1-carboxylate
687	3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)-N-[(3S)-1-piperidin-4-ylpyrrolidin-3-yl]pyrazine-2-carboxamide
688	1,1-dimethylethyl (2R)-2-[[[(3S)-3-[[[3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidin-1-yl]methyl]pyrrolidine-1-carboxylate
689	3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)-N-[(3S)-1-[(2R)-pyrrolidin-2-ylmethyl]pyrrolidin-3-yl]pyrazine-2-carboxamide
690	3-amino-N-[(3S)-1-[[4-(dimethylamino)phenyl]methyl]pyrrolidin-3-yl]-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
691	3-amino-N-[2-[[[4-(dimethylamino)phenyl]amino]carbonyl]amino]ethyl]-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
692	3-amino-N-[2-[[[4-(dimethylamino)phenyl]carbonyl]amino]ethyl]-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
693	3-{5-amino-6-[imino(2-pyridin-2-ylhydrazino)methyl]pyrazin-2-yl}-N-(phenylmethyl)benzamide
694	3-[5-amino-6-(morpholin-4-ylmethyl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
695	ethyl 1-[[3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]methyl]piperidine-4-carboxylate
696	3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)-N-[(3S)-1-(pyridin-2-ylmethyl)pyrrolidin-3-yl]pyrazine-2-carboxamide
697	methyl N-[[3-amino-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]threoninate
698	{3-[5-amino-6-(3-piperidin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
699	3-(ethylamino)-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
700	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-(3-[[[phenylmethyl]amino]carbonyl]phenyl)pyrazine-2-carboxamide
701	3-[5-amino-6-(1-methyl-5-piperidin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide

#	Name
702	3-[5-amino-6-(5-methyl-1-pyridin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
703	{3-[5-amino-6-(3-morpholin-2-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-yl]phenyl}methanol
704	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine
705	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-{3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-amine
706	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-ethylpyrrolidin-3-yl]pyrazine-2-carboxamide
707	6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-3-(ethylamino)-N-[(3S)-1-ethylpyrrolidin-3-yl]pyrazine-2-carboxamide
708	3-amino-6-[3-[[[(1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
709	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
710	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[[[(1R)-1-phenylethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
711	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-(3,4-dihydroisoquinolin-2(1H)-ylcarbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
712	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[[[(2-phenylethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
713	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[[[(1S)-1-phenylethyl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
714	3-amino-6-[3-(3,4-dihydroisoquinolin-2(1H)-ylcarbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
715	3-amino-6-(3-[[[(2-phenylethyl)amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
716	3-amino-6-[3-[[[(1R)-1-phenylethyl]amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
717	3-amino-6-[3-[[[(1S)-1-phenylethyl]amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
718	3-[5-amino-6-[(2-ethylhydrazino)(imino)methyl]pyrazin-2-yl]-N-(2,3-dihydro-1H-inden-1-yl)benzamide
719	3-[5-amino-6-[imino(2-methylhydrazino)methyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
720	3-[5-amino-6-[imino(2-phenylhydrazino)methyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
721	phenylmethyl 3-[3-[3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate
722	phenylmethyl 3-[3-[3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate
723	5-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]-3-(3-pyridin-4-yl-1H-1,2,4-triazol-5-yl)pyrazin-2-amine

#	Name
724	[3-(5-amino-6-{3-[(3R)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methanol
725	[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl}pyrazin-2-yl)phenyl]methanol
726	3-amino-6-{3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
727	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-[[[(1S,2R)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
728	1,1-dimethylethyl 3-[[[3-amino-6-{3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl}pyrazin-2-yl]carbonyl]amino]piperidine-1-carboxylate
729	3-amino-6-{3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]}-N-piperidin-3-ylpyrazine-2-carboxamide
730	3-amino-6-{3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]}-N-(1-ethylpiperidin-3-yl)pyrazine-2-carboxamide
731	3-amino-N-[(3S)-1-ethylpyrrolidin-3-yl]-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
732	(3-{5-amino-6-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]}pyrazin-2-yl)phenyl]methanol
733	5-{3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]}-3-[5-(1-ethylpiperidin-3-yl)-4H-1,2,4-triazol-3-yl]pyrazin-2-amine
734	3-[5-amino-6-(1-ethyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]benzamide
735	3-[5-amino-6-(1-methyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
736	(3-{5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]}pyrazin-2-yl)phenyl]methanol
737	5-{3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl]phenyl]}-3-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-amine
738	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
739	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl}pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
740	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[2-chloro-6-fluorophenyl]methyl]amino]carbonyl]phenyl}pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
741	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[[[4-(methylthio)phenyl]methyl]amino]carbonyl]phenyl}pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
742	1,1-dimethylethyl (3S)-3-[[[3-amino-6-{3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl}pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
743	3-amino-6-{3-[[[4-(methyloxy)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
744	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-[3-[[[4-[(trifluoromethyl)oxy]phenyl]methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
745	3-amino-6-{3-[(2,3-dihydro-1H-inden-2-ylamino)carbonyl]phenyl}-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide

#	Name
746	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
747	3-amino-6-[3-(((4-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
748	3-[5-amino-6-[imino(2-pyridin-4-ylhydrazino)methyl]pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]benzamide
749	phenylmethyl (3R)-3-[3-[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)pyrazin-2-yl]-1-ethyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate
750	3-amino-6-(3-[(1S)-1-[(1S)-2,3-dihydro-1H-inden-1-ylamino]ethyl]phenyl)-N-methylpyrazine-2-carboxamide
751	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide
752	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)pyrazin-2-yl]carbonyl]amino)methyl]pyrrolidine-1-carboxylate
753	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
754	phenylmethyl (3R)-3-[3-[3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)pyrazin-2-yl]-1-methyl-1H-1,2,4-triazol-5-yl]piperidine-1-carboxylate
755	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
756	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
757	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(pyridin-2-ylmethyl)amino]carbonyl]phenyl)pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
758	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[(pyridin-3-ylmethyl)amino]carbonyl]phenyl)pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
759	1,1-dimethylethyl (3S)-3-[[[3-amino-6-[3-[(3,4-dihydro-2H-chromen-4-ylamino)carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
760	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
761	3-amino-6-(3-[(pyridin-2-ylmethyl)amino]carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
762	3-amino-6-(3-[(pyridin-3-ylmethyl)amino]carbonyl]phenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
763	dimethyl 5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]benzene-1,3-dicarboxylate
764	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
765	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[(3R)-piperidin-3-yl]pyrazine-2-carboxamide
766	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-piperidin-3-ylpyrazine-2-carboxamide
767	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl)-N-[2-(1H-imidazol-4-yl)ethyl]pyrazine-2-carboxamide

#	Name
768	3-amino-N-azepan-4-yl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazine-2-carboxamide
769	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
770	1,1-dimethylethyl 4-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate
771	3-[5-amino-6-(5-{2-[(phenylmethyl)oxy]ethyl}-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
772	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)-4-fluorophenyl)-N-methylpyrazine-2-carboxamide
773	3-[5-amino-6-(1-phenyl-5-piperidin-3-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
774	3-[5-amino-6-[amino(imino)methyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
775	3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl)-N-[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]benzamide
776	1,1-dimethylethyl (3S)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
777	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate
778	3-amino-6-[3-[(3,4-dihydro-2H-chromen-4-ylamino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
779	3-amino-6-[3-[(2,3-dihydro-1-benzofuran-3-ylamino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
780	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[(3R)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
781	1,1-dimethylethyl 4-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)azepane-1-carboxylate
782	3-amino-N-azepan-4-yl-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
783	3-amino-N-azepan-4-yl-6-[3-[(3aR,8aS)-8,8a-dihydro-3aH-indeno[1,2-d][1,3]oxazol-2-yl]phenyl]pyrazine-2-carboxamide
784	1,1-dimethylethyl (3R)-3-(((3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
785	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)phenyl)-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
786	1,1-dimethylethyl (3R)-3-(((3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)methyl]pyrrolidine-1-carboxylate
787	3-amino-6-[3-(((1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl)amino)carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-ylmethyl]pyrazine-2-carboxamide
788	3-amino-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)carbonyl)-5-fluorophenyl)-N-methylpyrazine-2-carboxamide
789	3-(5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl)-5-[(methoxy)carbonyl]benzoic acid

#	Name
790	methyl 3-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-5-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]benzoate
791	5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]-N'-pyrrolidin-3-ylbenzene-1,3-dicarboxamide
792	3-[5-amino-6-(5-phenyl-1-pyridin-2-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-(phenylmethyl)benzamide
793	3-[5-amino-6-(5-methyl-1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
794	1,1-dimethylethyl (3R)-3-[[[3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino)piperidine-1-carboxylate
795	1,1-dimethylethyl (3R)-3-[[[3-amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino)piperidine-1-carboxylate
796	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
797	5-[5-amino-6-[(methylamino)carbonyl]pyrazin-2-yl]-N-[(1R)-2,3-dihydro-1H-inden-1-yl]-N'-[(3R)-pyrrolidin-3-yl]benzene-1,3-dicarboxamide
798	(3-[5-amino-6-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl]phenyl)methanol
799	3-[5-amino-6-(1,5-dimethyl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-[(1S)-2,3-dihydro-1H-inden-1-yl]benzamide
800	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[[[(1-methyl-1-phenylethyl)amino]carbonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
801	3-amino-6-(3-[[[(1-methyl-1-phenylethyl)amino]carbonyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
802	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl]phenyl]pyrazine-2-carboxamide
803	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl]phenyl]-N-piperidin-3-ylpyrazine-2-carboxamide
804	3-amino-6-(3-[[[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl]phenyl]-N-[(3S)-1-methylpyrrolidin-3-yl]pyrazine-2-carboxamide
805	3-amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[(3S)-1-methylpyrrolidin-3-yl]pyrazine-2-carboxamide
806	3-amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)methyl]phenyl]-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide
807	3-amino-6-[3-[[[(1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
808	3-amino-N-[(3S)-pyrrolidin-3-yl]-6-(3-[[[(1S)-1,2,3,4-tetrahydronaphthalen-1-ylamino]methyl]phenyl]pyrazine-2-carboxamide
809	N-[3-(5-amino-6-[5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl]pyrazin-2-yl]phenyl]-2-(2,4-dichlorophenyl)acetamide
810	1,1-dimethylethyl (3S)-3-[[[3-amino-6-(3-[[[(1R)-2,3-dihydro-1H-inden-1-ylamino]sulfonyl]phenyl]pyrazin-2-yl]carbonyl]amino]pyrrolidine-1-carboxylate
811	3-amino-6-(3-hydroxyphenyl)-N-[(3S)-pyrrolidin-3-yl]pyrazine-2-carboxamide

#	Name
812	3-amino-6-(3-(((1R)-2,3-dihydro-1H-inden-1-ylamino)sulfonyl)phenyl)-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
813	3-[5-amino-6-(1-pyridin-4-yl-1H-1,2,4-triazol-3-yl)pyrazin-2-yl]-N-((1S)-2,3-dihydro-1H-inden-1-yl)benzamide
814	3-amino-N-((3S)-pyrrolidin-3-yl)-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
815	3-amino-N-((3S)-pyrrolidin-3-yl)-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
816	3-amino-N-((3S)-pyrrolidin-3-yl)-6-[3-(((2,3,5-trifluorophenyl)methyl)amino)carbonyl)phenyl]pyrazine-2-carboxamide
817	3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl)phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
818	3-(5-amino-6-[3-(1-methylpiperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((2-chloro-6-fluorophenyl)methyl)benzamide
819	3-amino-N-1-azabicyclo[2.2.2]oct-3-yl-6-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)pyrazine-2-carboxamide
820	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((4-chloro-2-fluorophenyl)methyl)benzamide
821	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((2-chlorophenyl)methyl)benzamide
822	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
823	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]pyrazin-2-yl)carbonyl)amino)pyrrolidine-1-carboxylate
824	5-(3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl)-3-(4-piperidin-3-yl-1,3-thiazol-2-yl)pyrazin-2-amine
825	3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
826	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl)phenyl]-N-((3S)-pyrrolidin-3-yl)pyrazine-2-carboxamide
827	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((2,6-difluorophenyl)methyl)benzamide
828	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((2-chloro-6-fluorophenyl)methyl)benzamide
829	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((2,5-dichlorophenyl)methyl)benzamide
830	3-(5-amino-6-[3-((3S)-piperidin-3-yl)-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-((3,4-dichlorophenyl)methyl)benzamide
831	3-amino-6-[3-(((4-(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-((3S)-piperidin-3-yl)pyrazine-2-carboxamide
832	3-amino-6-[3-(((2,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-((3S)-piperidin-3-yl)pyrazine-2-carboxamide
833	1,1-dimethylethyl (3S)-3-(((3-amino-6-[3-(((1,1-dimethylethyl)oxy)carbonyl)amino)methyl)phenyl]pyrazin-2-yl)carbonyl)amino)piperidine-1-carboxylate

#	Name
834	3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(2-fluorophenyl)methyl]benzamide
835	3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(3-fluorophenyl)methyl]benzamide
836	3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(4-fluorophenyl)methyl]benzamide
837	3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(3-chlorophenyl)methyl]benzamide
838	3-(5-amino-6-{5-[(3S)-piperidin-3-yl]-4H-1,2,4-triazol-3-yl}pyrazin-2-yl)-N-[(4-chlorophenyl)methyl]benzamide
839	3-amino-6-[3-(aminomethyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
840	3-amino-6-(3-hydroxyphenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
841	1,1-dimethylethyl (3S)-3-[[[(3-amino-6-[3-(aminomethyl)phenyl]pyrazin-2-yl)carbonyl]amino]piperidine-1-carboxylate
842	3-amino-6-[3-[[[(4-fluorophenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
843	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[(4-(trifluoromethyl)oxy)phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
844	3-amino-6-[3-[[[biphenyl-4-yl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
845	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[(4-(1,2,3-thiadiazol-5-yl)phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
846	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[(2-(trifluoromethyl)oxy)phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
847	3-amino-6-[3-[[[biphenyl-2-yl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
848	3-amino-6-[3-[[[(2,4-difluorophenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
849	3-amino-6-[3-[[[(4-chloro-2-fluorophenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
850	3-amino-6-[3-[[[(4-fluoro-3-(trifluoromethyl)phenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
851	3-amino-6-[3-[[[naphthalen-2-yl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
852	3-amino-6-[3-[[[(4-(dimethylamino)phenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
853	3-amino-6-[3-[[[(1,3-benzodioxol-5-yl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
854	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[(4-(2-thienyl)phenyl)methyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
855	3-amino-6-[3-[[[(2-(methyloxy)phenyl)methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
856	3-amino-6-[3-(((2,4-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
857	3-amino-6-[3-(((4-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
858	3-amino-6-[3-(((2,3-difluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
859	3-amino-6-[3-(((4-(methylsulfonyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
860	3-amino-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-ylmethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
861	3-amino-6-[3-(((4-chlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
862	3-amino-6-[3-(((5-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
863	3-amino-N-[(3R)-1-azabicyclo[2.2.2]oct-3-yl]-6-[3-(((1S)-2,3-dihydro-1H-inden-1-ylamino)methyl)phenyl]pyrazine-2-carboxamide
864	3-amino-6-[3-(((4-(methylthio)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
865	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(2,4-difluorophenyl)methyl]benzamide
866	3-amino-6-[3-(((1,2-diphenylethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
867	3-(5-amino-6-[3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)-N-[(3,4-difluorophenyl)methyl]benzamide
868	3-amino-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
869	methyl 4-([3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl]pyrazin-2-yl)phenyl]carbamate
870	3-amino-6-[3-(((4-bromophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
871	3-amino-6-[3-(((4-bromo-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
872	3-amino-6-[3-(((2-bromophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
873	3-amino-6-[3-(((4-fluoro-2-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
874	3-amino-6-[3-(((2-fluoro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
875	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3R)-pyrrolidin-3-ylmethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
876	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3R)-pyrrolidin-3-ylamino)carbonyl]phenyl]pyrazine-2-carboxamide
877	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2-piperidin-1-ylethyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
878	3-amino-6-[3-(((2-phenylethyl)amino)carbonyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
879	3-amino-6-[3-(((2-(methoxy)ethyl)amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
880	3-amino-6-[3-(((4-iodophenyl)methyl)amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
881	3-amino-6-[3-(((2-chlorophenyl)methyl)amino)carbonyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
882	3-amino-6-[3-(((4-fluorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
883	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((4-(trifluoromethyl)oxy)phenyl)carbonyl)amino)methyl]phenyl]pyrazine-2-carboxamide
884	3-amino-6-[3-(((4-chlorophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
885	3-amino-6-(3-(((4-chlorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
886	3-amino-6-(3-(((2,6-dichlorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
887	3-amino-6-(3-(((pentafluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
888	3-amino-6-(3-(((2-chloro-4-fluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
889	3-amino-6-[3-(((4-(methylthio)phenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
890	3-amino-6-(3-(((2,4-dichlorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
891	3-amino-6-[3-(((4-(4-chlorophenyl)-2-thienyl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
892	3-amino-6-[3-(((4-bromophenyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
893	3-amino-6-(3-(((naphthalen-2-yl)carbonyl)amino)methyl)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
894	3-amino-6-(3-([2-(4-phenylpiperazin-1-yl)ethyl]oxy)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
895	N-[(3-{5-amino-6-[(3R)-piperidin-3-ylacetyl]pyrazin-2-yl}phenyl)methyl]-N'-(4-bromophenyl)urea
896	N-[(3-{5-amino-6-[(3R)-piperidin-3-ylacetyl]pyrazin-2-yl}phenyl)methyl]-N'-naphthalen-2-ylurea
897	3-amino-6-(3-(((2,4-difluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
898	3-amino-6-(3-(((2-fluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
899	3-amino-6-[3-(((5-[2-chloro-5-(trifluoromethyl)phenyl]furan-2-yl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
900	3-amino-6-[3-(((4-(dimethylamino)naphthalen-1-yl)carbonyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
901	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1,2,3-thiadiazol-5-yl)phenyl)carbonyl)amino)phenyl]pyrazine-2-carboxamide

#	Name
902	3-amino-6-(3-(((1,3-dimethyl-1H-pyrazol-5-yl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
903	3-amino-6-(3-(((4-(dimethylamino)phenyl)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
904	3-amino-6-(3-((2-hydroxyethyl)oxy)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
905	3-amino-6-(3-(((2,3-dihydro-1-benzofuran-5-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
906	3-amino-6-(3-(((3,4-dihydro-2H-1,5-benzodioxepin-7-ylsulfonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
907	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((quinolin-7-ylsulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
908	3-amino-6-(3-(((biphenyl-4-ylamino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
909	3-amino-6-(3-(((4-bromo-2-fluorophenyl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
910	N-[3-(5-amino-6-(((3S)-piperidin-3-ylamino)carbonyl)pyrazin-2-yl)phenyl]quinoline-3-carboxamide
911	3-amino-6-(3-(((6-(methyloxy)-1-benzofuran-3-yl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
912	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(trifluoromethyl)biphenyl-2-yl)carbonyl)amino)phenyl]pyrazine-2-carboxamide
913	3-amino-6-(3-(((3,5-bis(trifluoromethyl)phenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
914	3-amino-6-(3-((((2,4-difluorophenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
915	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2-(trifluoromethyl)phenyl)sulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
916	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((4-((trifluoromethyl)oxy)phenyl)sulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
917	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-(((2-((trifluoromethyl)oxy)phenyl)sulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
918	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-((((4-(trifluoromethyl)phenyl)sulfonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
919	3-amino-N-[(3S)-piperidin-3-yl]-6-(3-((((2-(trifluoromethyl)oxy)phenyl)amino)carbonyl)amino)methyl)phenyl)pyrazine-2-carboxamide
920	3-amino-6-(3-(((2-(methylthio)phenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
921	3-amino-6-(3-((((3-bromo-5-methyl)phenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
922	3-amino-6-(3-((((2-bromophenyl)amino)carbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
923	5-[3-(aminomethyl)phenyl]-3-(3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl)pyrazin-2-amine

#	Name
924	3-amino-6-{3-[[[4-(4-methylpiperazin-1-yl)phenyl]methyl]amino]carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
925	3-amino-6-{3-[[[2-(4-fluorophenyl)ethyl]amino]carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
926	3-amino-6-{3-[[[2-morpholin-4-ylethyl]amino]carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
927	3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[[[2R]-pyrrolidin-2-ylmethyl]amino]carbonyl]phenyl]pyrazine-2-carboxamide
928	3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[[[2S]-pyrrolidin-2-ylamino]carbonyl]phenyl]pyrazine-2-carboxamide
929	3-amino-6-{3-[(cyclopentylamino)carbonyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
930	3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[[[4-[(trifluoromethyl)oxy]phenyl]amino]carbonyl]amino]methyl]phenyl]pyrazine-2-carboxamide
931	3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[[[4-(trifluoromethyl)phenyl]amino]carbonyl]amino]methyl]phenyl]pyrazine-2-carboxamide
932	3-amino-6-{3-[[[4-(methyloxy)phenyl]carbonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
933	3-amino-6-{3-[[[biphenyl-4-ylcarbonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
934	3-amino-6-{3-[[[furan-2-ylcarbonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
935	3-amino-6-{3-[[[2,3-dihydro-1,4-benzodioxin-6-ylamino]carbonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
936	3-amino-6-{3-[[[2,3-dihydro-1-benzofuran-5-ylcarbonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
937	3-amino-6-{3-[[[2,3-dihydro-1,4-benzodioxin-6-ylsulfonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
938	3-amino-N-[(3S)-piperidin-3-yl]-6-{3-[[[2-thienylcarbonyl]amino]methyl]phenyl]pyrazine-2-carboxamide
939	3-amino-6-{3-[[[2,3-dihydro-1,4-benzodioxin-2-ylcarbonyl]amino]methyl]phenyl}-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
940	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)phenyl]methyl]-2-chlorobenzamide
941	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)phenyl]methyl]-4-chlorobenzamide
942	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)phenyl]methyl]-4-iodobenzamide
943	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)phenyl]methyl]-3,5-difluorobenzamide
944	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)phenyl]methyl]-4-bromo-2-fluorobenzamide
945	N-[[3-(5-amino-6-{3-[(3S)-piperidin-3-yl]-1H-1,2,4-triazol-5-yl]pyrazin-2-yl)phenyl]methyl]-4-bromo-2-chlorobenzamide

#	Name
946	3-[3-(2-aminopyridin-4-yl)-1H-1,2,4-triazol-5-yl]-5-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenylpyrazin-2-amine
947	3-amino-N-azepan-3-yl-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]methyl)phenylpyrazine-2-carboxamide
948	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[2-(2-thienyl)-1,3-thiazol-4-yl]carbonyl]amino]methyl]phenylpyrazine-2-carboxamide
949	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[5-(2-thienyl)pyridin-3-yl]carbonyl]amino]methyl]phenylpyrazine-2-carboxamide
950	3-amino-6-[3-[[[4-(1,1-dimethylethyl)phenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
951	3-amino-6-[3-[[[2,4-dichloro-6-methylphenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
952	3-amino-6-[3-[[[3,5-dichlorophenyl]methyl]amino]carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
953	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[2-thienylsulfonyl]amino]methyl]phenyl]pyrazine-2-carboxamide
954	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[3-thienylsulfonyl]amino]methyl]phenyl]pyrazine-2-carboxamide
955	3-amino-6-[3-[[[4-chlorophenyl]sulfonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
956	3-amino-6-[3-[[[1,3-benzodioxol-5-ylamino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
957	3-amino-6-[3-[[[4-(methyloxy)phenyl]amino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
958	3-amino-6-[3-[[[4-fluorophenyl]amino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
959	3-amino-6-[3-[[2-(diethylamino)ethyl]oxy]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
960	3-amino-6-[3-[[2-(morpholin-4-ylethyl)oxy]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
961	3-amino-6-[3-(methyloxy)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
962	3-amino-6-[3-(ethyloxy)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
963	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[[[4-(1,2,3-thiadiazol-4-yl)phenyl]methyl]amino]carbonyl]phenylpyrazine-2-carboxamide
964	3-amino-6-[3-[[2-(ethylamino)ethyl]oxy]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
965	phenylmethyl 4-(2-[[3-(5-amino-6-[(3S)-piperidin-3-ylamino]carbonyl]pyrazin-2-yl)phenyl]oxy)ethyl)piperazine-1-carboxylate
966	3-amino-6-[3-[[[4-(dimethylamino)phenyl]amino]carbonyl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
967	3-amino-N-azepan-3-yl-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenylpyrazine-2-carboxamide
968	3-amino-6-[3-[[[1R,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino]methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
969	3-amino-6-[3-(((1S,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl)phenyl]-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide
970	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((1S,2S)-2-hydroxy-2,3-dihydro-1H-inden-1-yl]amino)carbonyl)phenyl]pyrazine-2-carboxamide
971	3-amino-6-[3-(((4-chlorophenyl)amino)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
972	3-amino-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl)-N-[(3S)-1-(phenylmethyl)azepan-3-yl]pyrazine-2-carboxamide
973	3-amino-N-[(3S)-azepan-3-yl]-6-(3-[(1S)-2,3-dihydro-1H-inden-1-ylamino]carbonyl)phenyl]pyrazine-2-carboxamide
974	3-amino-N-[(3S)-azepan-3-yl]-6-(3-methylphenyl)pyrazine-2-carboxamide
975	3-amino-6-(3-[(2-methylphenyl)acetyl]amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
976	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3-(trifluoromethyl)phenyl)acetyl)amino)phenyl]pyrazine-2-carboxamide
977	3-amino-6-[3-(((4-(methyloxy)phenyl)acetyl)amino)phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
978	3-amino-6-(3-(((6-methylpyridin-3-yl)carbonyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
979	3-amino-6-(3-(((3-fluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
980	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[(pyridin-3-ylacetyl)amino]phenyl]pyrazine-2-carboxamide
981	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-[(pyridin-3-ylcarbonyl)amino]phenyl]pyrazine-2-carboxamide
982	3-amino-6-(3-(((2,5-difluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
983	3-amino-6-(3-(((4-fluorophenyl)acetyl)amino)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
984	3-amino-6-[3-(((3-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
985	3-amino-6-[3-(((3,4-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
986	3-Amino-6-[3-(((5-(4-nitro-phenyl)-furan-2-carbonyl)-amino)-methyl)-phenyl]-pyrazine-2-carboxylic acid (3S)-piperidin-3-ylamide
987	3-amino-6-[3-(((3-hydroxypyridin-2-yl)carbonyl)amino)methyl]phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
988	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(trifluoromethyl)oxy)phenyl)oxy)acetyl]amino)methyl]phenyl]pyrazine-2-carboxamide
989	3-amino-6-[3-(((5-methylisoxazol-3-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
990	3-amino-6-(3-(((isoxazol-5-ylcarbonyl)amino)methyl)phenyl)-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
991	3-amino-6-[3-(((4-methyl-1,2,3-thiadiazol-5-yl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
992	3-amino-6-[3-(((2,5-dichloro-3-thienyl)carbonyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
993	3-amino-6-[3-(((5-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
994	3-amino-6-[3-(((2-chloro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
995	3-amino-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
996	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((4-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
997	3-amino-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
998	3-amino-6-[3-(((2-fluoro-5-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
999	3-amino-6-[3-(((2,4-difluorophenyl)methyl)amino)methyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1000	3-amino-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1001	3-amino-6-[3-(((2-fluoro-4-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1002	3-amino-6-[3-(((2-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1003	3-amino-6-[3-(((2,3-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1004	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,6-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1005	3-amino-6-[3-(((4-chloro-3-(trifluoromethyl)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1006	3-amino-6-[3-(((3-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1007	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1008	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1009	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3,5-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1010	3-amino-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1011	3-amino-6-[3-(((3,4-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1012	3-amino-6-[3-(((6-chloro-2-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
1013	3-amino-6-[3-(((2-chloro-6-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1014	3-amino-6-[3-(((2-chloro-3,6-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1015	3-amino-6-[3-(((2,5-dichlorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1016	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,4-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1017	3-amino-6-[3-(((2,3-difluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1018	3-amino-6-[3-(((3-fluoro-4-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1019	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1020	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,3-dihydro-1,4-benzodioxin-6-yl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1021	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2-chloro-6-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1022	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((2,6-difluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1023	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-4-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1024	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((3-chloro-2-fluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1025	3-amino-6-[3-(((3-chloro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1026	3-amino-6-[3-(((2,6-bis(methoxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1027	3-amino-6-[3-(((3-bromo-4-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1028	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((3,4,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1029	3-amino-6-[3-(((4-fluoro-3-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1030	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((2,3,5-trifluorophenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1031	3-amino-6-[3-(((5-fluoro-2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1032	3-amino-6-[3-(((3-bromo-5-fluorophenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1033	3-amino-N-[(3S)-piperidin-3-yl]-6-[3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide
1034	3-amino-6-[3-(((biphenyl-3-yl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1035	3-amino-6-[3-(((2-methylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide

#	Name
1036	3-amino-6-[3-(((2,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1037	3-amino-6-[3-(((3,5-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1038	3-amino-6-[3-(((2,4-dimethylphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1039	3-amino-6-[3-(((3-methoxyphenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1040	3-amino-6-[3-(((phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1041	3-amino-6-[3-(((3,5-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1042	3-amino-6-[3-(((3,4-bis(methyloxy)phenyl)methyl)amino)carbonyl]phenyl]-N-[(3S)-piperidin-3-yl]pyrazine-2-carboxamide
1043	3-amino-N-[(3S)-azepan-3-yl]-6-[3-(((4-(1H-pyrazol-1-yl)phenyl)methyl)amino)carbonyl]phenyl]pyrazine-2-carboxamide

43. A pharmaceutical composition comprising a compound according to any one of claims 1 - 42 and a pharmaceutically acceptable carrier.

44. A metabolite of the compound or the pharmaceutical composition according to any one of claims 1 - 43.

45. A method of modulating the *in vivo* activity of a kinase, the method comprising administering to a subject a therapeutically effective amount of the compound or the pharmaceutical composition according to any of claims 1 - 43 to a mammal in need thereof.

46. The method according to claim 45, wherein the kinase is Chk1 or Chk2.

47. The method according to claim 45, wherein the kinase is Chk1.

48. The method according to claim 46, wherein modulating the *in vivo* activity of the kinase comprises inhibition of said kinase.

49. A method of inhibiting proliferation of a plurality of cancerous cells, the method comprising contacting the plurality of cancerous cells, or causing the plurality of cancerous cells to be contacted with, a cancer therapeutic and the compound as described in any one of claims 1 - 42.

50. The method according to claim 49, wherein the cancer therapeutic is a DNA damaging agent.

51. A method of diminishing proliferation of a plurality of cancerous cells in a patient, the method comprising administering to the patient the compound as described in any one of claims 1 – 42, and one or more cancer therapeutic agents.

52. The method of claim 51, wherein the compound is combined with a pharmaceutically acceptable carrier.

53. The method of claim 52, wherein the cancer therapeutic agent is selected from the group consisting of a Topoisomerase I inhibitor, a Topoisomerase II inhibitor, an alkylating agent, a nitrogen mustard, an aziridine, an epoxide, an alkyl sulfonate, a nitrosourea, an alkylating agent-steroid conjugate, a DNA damaging/binding agent, a Cisplatin, an antibiotic, an anthracycline, an anthracenedione, an antimetabolite, an antifolate, a nucleic acid analog, a ribonucleic acid analog, a ribozyme, radiation, a Vinca alkaloid, a Taxane, an enzyme, a natural product, a kinase inhibitor, a hormone a hormones antagonist, and a biological response modifier.

54. The method of claim 53, wherein where the cancer therapeutic agent comprises at least one of: Camptothecin, Topotecan, 9-Nitrocamptothecin, 9-Aminocamptothecin, Karenitecin, Irinotecan, Etoposide, Etoposide Phosphate, Teniposide, Amsacrine, Razoxane, Dexrazoxane, Mechlorethamine, Cyclophosphamide, Ifosfamide, Chlorambucil, Melphalan, Thiotepa, Trenimon, Triethylenemelamine, Dianhydrogalactitol, Dibromodulcitol, Busulfan, dimethylsulfate, Chloroethylnitrosourea, BCNU, CCNU, Methyl-CCNU, Streptozotocin, Chlorozotocin, Prednimustine, Estramustine, Procarbazine, Dacarbazine, Hexamethylmelamine, Pentamethylmelamine, Temozolomide, Cisplatin, Carboplatin, Oxaliplatin, Bleomycin, Dactinomycin, Mithramycin, Mitomycin C, Daunorubicin, Doxorubicin, Epirubicin, Idarubicin, Methotrexate, Edatrexate, Trimethoprim, Nolatrexed, Raltitrexed, Hydroxyurea, 5-fluorouracil, Ftorafur, Capecitabine, Furtulon, Eniluracil, ara-C, 5-azacytidine, Gemcitabine, Mercaptopurine, Thioguanine, Pentostatin, antisense DNA, antisense RNA, an antisense DNA/RNA hybrid, a ribozyme, ultraviolet radiation, Vincristine, Vinblastine, Paclitaxel, Docetaxel, L-Asparaginase, a kinase inhibitor, Imatinib, Mitotane, Aminoglutethimide, Diethylstilbestrol, Ethinyl estradiol, Tamoxifen, Anastrozole, Testosterone propionate, Fluoxymesterone, Flutamide, Leuprolide, Prednisone, Hydroxyprogesterone caproate, Medroxyprogesterone acetate, Megestrol acetate, Interferon-alfa, and Interleukin.

55. A method of enhancing the efficacy of a disease therapy that inflicts DNA damage, the method comprising combining the disease therapy with administration of a compound or pharmaceutical composition as described in any one of claims 1 - 43.

56. A method of treating a kinase-dependent disease or condition, the method comprising administering a therapeutically effective amount of the compound according to any one of claims 1 - 43 to a mammal in need thereof.

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(71) Applicant (for all designated States except US): **EX-ELIXIS, INC.** [US/US]; P.O. Box 511, 170 Harbor Way, South San Francisco, CA 94083-0511 (US).

(72) Inventors; and

(75) Inventors/Applicants (for US only): **BUHR, Chris, A.** [US/US]; 355 Belmont Avenue, Redwood City, CA 94061 (US). **BAIK, Tae-Gon** [KR/US]; 1125 Balclutha Drive, #105, Foster City, CA 94404 (US). **MA, Sunghoon** [KR/US]; 1110 Polynesia Drive, #109, Foster City, CA 94404 (US). **TESFAI, Zerom** [ER/US]; 4704 Springlake Drive, San Leandro, CA 94578 (US). **WANG, Longcheng** [CN/US]; 22 Skypark Circle, South San Francisco, CA 94080 (US). **CO, Erick, Wang** [US/US]; 921 Salt Court, Redwood City, CA 94065 (US). **EPSHTEYN, Sergey** [US/US]; 2922 Southwycke Terrace, Fremont, CA 94536 (US). **KENNEDY, Abigail, R.** [US/US]; 325 West Lowell Avenue, Tracy, CA 95376 (US). **CHEN, Baili** [US/US]; 1054 Colorado Avenue, Palo Alto, CA 94303 (US). **DUBENKO, Larisa** [US/US]; 3326 Balboa Street, San Francisco, CA 94121 (US). **ANAND, Neel Kumar** [GB/US]; 2713 South Norfolk Street, Apt.#206, San Mateo, CA 94403 (US). **TSANG, Tsze, H.** [US/US]; 8367 Kent Drive, El Cerrito, CA 94530 (US). **NUSS, John, M.** [US/US]; 16 Woodranch Circle, Danville, CA 94056 (US). **PETO, Csaba, J.** [HU/US]; 965 Shorepoint Court, #305, Alameda, CA 94501 (US). **RICE, Kenneth, D.** [US/US]; 802 Autumn Lane, Mill Valley, CA 94941 (US). **IBRAHIM, Mohamed, Abdulkader** [US/US]; 3380 Lubich Drive, Mountain View, CA 94040 (US). **SCHNEPP, Kevin, Luke** [US/US]; 8768 Newry Court, Elk Grove,

CA 95624 (US). **SHI, Xian** [CN/US]; 157 Elm Street, Apt. 106, San Mateo, CA 94401 (US). **LEAHY, James, William** [US/US]; 1185 Camellia Court, San Leandro, CA 94577 (US). **CHEN, Jeff** [US/US]; 417 Fulton Street #6, San Francisco, CA 94102 (US). **DALRYMPLE, Lisa, Esther** [US/US]; 387 17th Avenue, San Francisco, CA 94121 (US). **FORSYTH, Thlmothy, Patrick** [US/US]; 2071A Isabelle Avenue, San Mateo, CA 94403 (US). **HUYNH, Tai, Phat** [US/US]; 1530 6th Avenue, #2, Oakland, CA 94606 (US). **MANN, Grace** [US/US]; 231 Callippe Court, Brisbane, Ca 94005 (US). **MANN, Lary, Wayne** [US/US]; 780 Bair Island Road, Apt. 304, Redwood City, CA 94063 (US). **TAKEUCHI, Craig, Stacy** [CA/US]; 1090 Carolan Avenue, #302, Burlingame, CA 94010 (US).

(74) Agents: **GRIEDEL, Brian et al.**; Exelixis, Inc., P.O. Box 511, 170 Harbor Way, South San Francisco, CA 94083-0511 (US).

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For two-letter codes and other abbreviations, refer to the "Guidance Notes on Codes and Abbreviations" appearing at the beginning of each regular issue of the PCT Gazette.

(54) Title: PROTEIN KINASE MODULATORS AND METHODS OF USE

(57) Abstract: This invention relates to compounds for modulating protein kinase enzymatic activity for modulating cellular activities such as proliferation, differentiation, programmed cell death, migration and chemoinvasion, and to pharmaceutical compositions containing such compounds. Even more specifically, the invention relates to compounds that inhibit, regulate and/or modulate kinases, particularly Checkpoint Kinases, even more particularly Checkpoint Kinase 1, or Chk1. Methods of therapeutically or prophylactically using the compounds and compositions to treat kinase-dependent diseases and conditions are also an aspect of the invention, and include methods of treating cancer, as well as other disease states associated with unwanted angiogenesis and/or cellular proliferation, by administering effective amounts of such compounds.

WO 2003/093297 A3

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/13869

A. CLASSIFICATION OF SUBJECT MATTER

IPC(7) : A61K 31/4965, 31/497, 31/55; C07D 241/28, 401/04, 401/06, 401/12, 405/04, 413/04, 413/14.
US CL : 514/217.05, 252.11, 255.05, 255.06; 540/598; 544/357, 405, 407.

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 514/217.05, 252.11, 255.05, 255.06; 540/598; 544/357, 405, 407.

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)
CAS ONLINE STRUCTURE SEARCH

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	WO 00/12488 A1 (GLAXO GROUP LIMITED) 09 March 2000 (09.03.2000), see especially species on pages 4-5.	1,2,43
X	WO 97/43267 A1 (KINGS COLLEGE) 20 November 1997 (20.11.1997), see especially pages 14,17-18.	1-5,21-22

☐ Further documents are listed in the continuation of Box C.

☐ See patent family annex.

* Special categories of cited documents:	* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"A" document defining the general state of the art which is not considered to be of particular relevance	"X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"E" earlier application or patent published on or after the international filing date	"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"&" document member of the same patent family
"O" document referring to an oral disclosure, use, exhibition or other means	
"P" document published prior to the international filing date but later than the priority date claimed	

Date of the actual completion of the international search

09 February 2004 (09.02.2004)

Date of mailing of the international search report

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Mail Stop PCT, Attn: ISA/US
Commissioner for Patents
P.O. Box 1450
Alexandria, Virginia 22313-1450

Facsimile No. (703) 305-3230

Authorized officer

Valerie Bell-transfr

Telephone No. 571/272-1600

INTERNATIONAL SEARCH REPORT

International application No.

PCT/US03/13869

Box I Observations where certain claims were found unsearchable (Continuation of Item 1 of first sheet)

This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:

1. ☐ Claim Nos.:
because they relate to subject matter not required to be searched by this Authority, namely:
2. ☒ Claim Nos.: 1-41 (in part), 43 (in part), 44, 49-56 (in part)
because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:
Please See Continuation Sheet
3. ☒ Claim Nos.: 45-48
because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).

Box II Observations where unity of invention is lacking (Continuation of Item 2 of first sheet)

This International Searching Authority found multiple inventions in this international application, as follows:

1. ☐ As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.
 2. ☐ As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.
 3. ☐ As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos.:
 4. ☐ No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:
- Remark on Protest ☐ The additional search fees were accompanied by the applicant's protest.
☐ No protest accompanied the payment of additional search fees.

INTERNATIONAL SEARCH REPORT

PCT/US03/13869

Continuation of Box I Reason 2:

The present claims fail to comply with the clarity, conciseness, support and disclosure requirements of PCT Articles 5 and 6 to such an extent that a meaningful search for the full scope of the claims is impossible. Consequently, the search has been carried out for compounds specifically described in claim 42. The scope of "metabolite" in claim 44 is not all ascertainable since description provides no guidance as to structural makeup.